MM 18 25

appropriate serial number.

Scientific and Technical Information Center

SEARCH REQUEST FORM

Requester's Full Name: Koy P. TSSac Examiner #: 82353 Date: 45/18/06
Art Unit: 1623 Phone Number: 2- 2674 Serial Number: 10/654365
Location (Bldg/Room#): 5D24 (Mailbox #): 5D24 Results Format Preferred (circle): PAPER DISK or Email

To ensure an efficient and quality search, please attach a copy of the cover sheet, claims, and abstract or fill out the following:
Title of Invention: TREATMENT OF NON-ALLERGIC RHINITIS BY SELECTIVE PHOSPHO DIESTERASE-4
Inventors (please provide full names): RUNDELDT C.
RELIGE HILDEGARD, K: HOFGEN, N.
Earliest Priority Date: 06 September 2002
Search Topic: Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc., if known.
For Sequence Searches Only Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the

Type of Search Vendors and cost where applicable NA Sequence (#) Questel/Orbit _Lexis/Nexis AA Sequence (#) Westlaw WWW/Internet Date Searcher Picked Up: 5/19/06 _In-house sequence systems Date Completed: 5/23/06 Commercial Score/Length SPDI Encode/Transl Searcher Prep & Review Time: 45 Fulltext Online Time:

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             51) SEA ABB=ON PLU=ON ("KUSS H"/AU OR "KUSS H H"/AU OR "KUSS H
                J"/AU OR "KUSS H M"/AU OR "KUSS HILDEGARD"/AU)
             12) SEA ABB=ON PLU=ON ("HOFGEN N"/AU OR "HOFGEN NORBERT"/AU)
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L4
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                ACTIVATE KUSSALL/A
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L5
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                J"/AU OR "KUSS H M"/AU OR "KUSS HILDEGARD"/AU)
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                D L9
              0 SEA SSS SAM L9
L11
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L12
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                D SCAN
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L17
                QUE ABB=ON PLU=ON L16
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2 SEA SSS SAM L16

L18

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D SCAN
L19 35 SEA SSS FUL L16
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FILE 'CAPLUS' ENTERED AT. 11:22:00 ON 23 MAY 2006 51 SEA ABB=ON PLU=ON L19 L20 10 SEA ABB=ON PLU=ON L20 NOT (PY>2002 OR AY>2002 OR PRY >2002) L21 11 SEA ABB=ON PLU=ON L22L20 AND L8 19 SEA ABB=ON PLU=ON L23 (L22 OR L21) L24 14 SEA ABB=ON PLU=ON (L4 OR L22) 40 SEA ABB=ON PLU=ON L20 NOT L22 L25 FILE 'REGISTRY' ENTERED AT 11:24:26 ON 23 MAY 2006 L26 ANALYZE PLU=ON L19 1-35 RN : D

FILE 'CAPLUS' ENTERED AT 11:26:38 ON.23 MAY 2006 L30 10 SEA ABB=ON PLU=ON L29

=> file caplus FILE 'CAPLUS' ENTERED AT 11:28:38 ON 23 MAY 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 23 May 2006 VOL 144 ISS 22 FILE LAST UPDATED: 22 May 2006 (20060522/ED)

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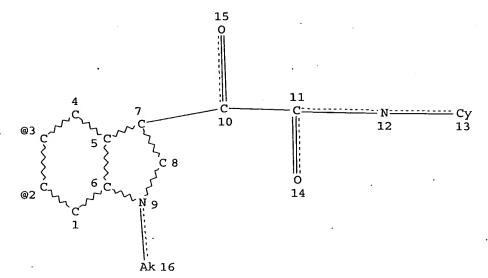
http://www.cas.org/infopolicy.html
'OBI' IS DEFAULT SEARCH FIELD FOR 'CAPLUS' FILE

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L6 ( 51)SEA FILE=CAPLUS ABB=ON PLU=ON ("KUSS H"/AU OR "KUSS H H"/AU OR "KUSS H J"/AU OR "KUSS H M"/AU OR "KUSS HILDEGARD"/AU)
L7 ( 12)SEA FILE=CAPLUS ABB=ON PLU=ON ("HOFGEN N"/AU OR "HOFGEN
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Saloni Sharma 05/23/2006

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NORBERT"/AU)
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L8 126 SEA FILE=CAPLUS ABB=ON PLU=ON (L5 OR L6 OR L7) L16 STR



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VPA 17-2/3 U

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GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 17

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DEFAULT ECLEVEL IS LIMITED

STEREO ATTRIBUTES: NONE

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L25	40	SEA	FILE=CAPLUS	ABB=ON	PLU=ON	L20	NOT	L22

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=> d ibib abs hitstr 125 tot
L25 ANSWER 1 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                         2006:365169 CAPLUS
DOCUMENT NUMBER:
                         144:419682
TITLE:
                         Pharmaceutical compositions containing
                         phosphodiesterase IV inhibitors and immunosuppressants
INVENTOR(S):
                         Harada, Daisuke; Kobayashi, Katsuya; Manabe, Haruhiko;
                         Ohshima, Etsuo
PATENT ASSIGNEE(S):
                         Kyowa Hakko Kogyo Co., Ltd., Japan
SOURCE:
                         PCT Int. Appl., 78 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         Japanese
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                         KIND
                               DATE
                                           APPLICATION NO.
                                                                  DATE
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     WO 2006041120
                                20060420
                                           WO 2005-JP18854
                         A1
                                                                  20051013
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ,
             NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG,
             SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN,
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         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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             KG, KZ, MD, RU, TJ, TM
PRIORITY APPLN. INFO.:
                                           JP 2004-299104
                                                               A 20041013
                                           JP 2005-113265
                                                               A 20050411
     This invention relates to pharmaceutical compns. for the prevention and
AB
```

treatment of chronic skin diseases, comprising (a) a phosphodiesterase

(PDE)-IV inhibitor or a pharmacol. acceptable salt thereof and (b) an immunosuppressant, which are administered simultaneously or sep. with an interval. For example, tablets were formulated containing 2-(3,5-dichloro-4-pyridinyl)-1-(7-methoxyspiro[1,3-benzodioxole-2,1'cyclopentan]-4-yl)ethanone (PDE-IV inhibitor) 20, tacrolimus (immunosuppressant) 20, lactose 123.4, starch 20, hydroxypropyl cellulose 6, and Mg stearate 0.6 mg per tablet.

IT 257892-33-4

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(phosphodiesterase IV inhibitor and immunosuppressant combinations for treatment of chronic skin diseases)

257892-33-4 CAPLUS RN

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

Saloni Sharma 05/23/2006

REFERENCE COUNT:

113 THERE ARE 113 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 2 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2006:364924 CAPLUS

DOCUMENT NUMBER:

144:398341

TITLE:

Phosphodiesterase IV inhibitor and steroid

combinations for the treatment of chronic skin disease Harada, Daisuke; Kobayashi, Katsuya; Manabe, Haruhiko;

INVENTOR(S): Harada, Daisuk Ohshima, Etsuo

PATENT ASSIGNEE(S):

Kyowa Hakko Kogyo Co., Ltd., Japan

SOURCE:

PCT Int. Appl., 67 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATI	PATENT NO.				KIN	D :	DATE		i	APPL	ICAT	ION I	NO.		Di	ATE	
WO 2	2006	 0411	21.		Δ1	-	 2006:	0420	1	 ₩Ω 2	 005-	TD18	 855		- ·	0051	012
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PRIORITY APPLN. INFO.:

JP 2004-299103 .A 20041013 JP 2005-113264 A 20050411

AB It is intended to provide a remedy and/or a preventive for a chronic skin disease which comprises (a) a phosphodiesterase (PDE)-IV inhibitor or a pharmacol. acceptable salt thereof and (b) a steroid drug, which are administered simultaneously or sep. at an interval. For example, tablets were formulated containing 2-(3,5-dichloro-4-pyridinyl)-1-(7-methoxyspiro[1,3-benzodioxole-2,1'-cyclopentan]-4-yl)ethanone 50, prednisolone 20, lactose

123.4, starch 20, hydroxypropyl cellulose 6, and Mg stearate 0.6 mg per tablet.

IT 257892-33-4, AWD 12-281

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Usés)

(phosphodiesterase IV inhibitor and steroid combinations for treatment of chronic skin disease)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & &$$

REFERENCE COUNT:

128 THERE ARE 128 CITED REFERENCES AVAILABLE FOR

THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L25 ANSWER 3 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2006:149262 CAPLUS

DOCUMENT NUMBER:

144:239931

TITLE:

Pharmaceutical compositions for the treatment of

respiratory and gastrointestinal disorders

INVENTOR (S):

Jung, Birgit; Himmelsbach, Frank

PATENT ASSIGNEE(S):

Boehringer Ingelheim International GmbH, Germany;

Boehringer Ingelheim Pharma Gmbh & Co. KG

SOURCE:

PCT Int. Appl., 321 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE: FAMILY ACC. NUM. COUNT:

PA	TENT 1	NO.			KIN	D 1	DATE		i	APPL:	ICAT:	ION I	NO.		D	ATE	
						-		÷									
WO	2006	0157	75	•	A2	:	2006	0216	1	WO 2	005-1	EP83	85		20	00508	303
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IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

US 2006035893 A1 20060216 US 2005-189643 20050726 PRIORITY APPLN. INFO.: EP 2004-18808 Α 20040807 OTHER SOURCE(S): MARPAT 144:239931

The present invention relates to novel pharmaceutical compns. comprising at least 1 EGFR kinase inhibitor and at least one addnl. active compound selected from β -2 mimetics, steroids, PDE-IV inhibitors, p38 MAP kinase inhibitors, NK1 antagonists and endothelin-antagonists, processes for preparing the compns. and the use thereof as drugs in the treatment of respiratory or gastrointestinal complaints, as well as inflammatory diseases of the joints, the skin or the eyes. Thus, an inhalable powder contained an EGFR kinase inhibitor 150, formoterol fumarate dihydrate 50, and lactose 12,300 mg/capsule.

257892-33-4 AWD 12-281

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical compns. for treatment of respiratory and gastrointestinal disorders)

RN257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

L25 ANSWER 4 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:46803

DOCUMENT NUMBER: 144:135233

TITLE: Pharmaceuticals for inhalation comprising PDE IV

inhibitors and glycopyrrolate salts

CAPLUS

PATENT ASSIGNEE(S): Boehringer Ingelheim Pharma Gm.b.H. & Co. K.-G.,

Germany

SOURCE: Eur. Pat. Appl., 24 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

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EP 1616567
                           20060118
                                       EP 2004-16878
                                                               20040716
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        IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR
                     A1
                           20060126
                                      WO 2005-EP52704
                                                               20050613
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        CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
        GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
        LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
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        ZA, ZM, ZW
    RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
        IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF,
        CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM,
        KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG,
        KZ, MD, RU, TJ, TM
                                       EP 2004-16878
                                                           A 20040716
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PRIORITY APPLN. INFO.:

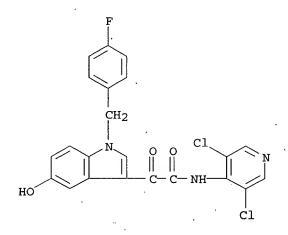
The present invention relates to novel pharmaceutical compns. based on PDE IV inhibitors and salts of glycopyrrolate salts, processes for preparing them and their use in the treatment of respiratory complaints. Thus, a formulation contained a glycopyrrolate salt 60, AWD 12281 200, lactose

12240 µg/capsule. IT **257892-33-4**, AWD 12-281

> RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (GW 842470; pharmaceuticals for inhalation comprising PDE IV inhibitors and glycopyrrolate salts)

RN257892-33-4 CAPLUS

CN1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2006 ACS on STN L25 ANSWER 5 OF 40

3

ACCESSION NUMBER: 2005:1289733 CAPLUS

DOCUMENT NUMBER: 144:40794

TITLE: Combinations comprising antimuscarinic agents and PDE4

inhibitors

INVENTOR(S): Gras Escardo, Jordi; Llenas Calvo, Jesus; Ryder,

Hamish; Orviz Diaz, Pio

PATENT ASSIGNEE(S): Almirall Prodesfarma S.A., Spain SOURCE:

PCT Int. Appl., 43 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

	PATENT NO.		APPLICATION NO.	
	WO 2005115465	71 20051200	WO 2005-EP5839	
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			MA, MD, MG, MK, MN,	
			PL, PT, RO, RU, SC,	
	SL, SM, SY,	TJ. TM. TN. TR.	TT, TZ, UA, UG, US,	UZ. VC. VN. VII
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	RW: BW, GH, GM,	KE, LS, MW, MZ,	NA, SD, SL, SZ, TZ,	UG, ZM, ZW, AM.
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	EE, ES, FI,	FR, GB, GR, HU,	IE, IS, IT, LT, LU,	MC, NL, PL, PT,
	RO, SE, SI,	SK, TR, BF, BJ,	CF, CG, CI, CM, GA,	GN, GQ, GW, ML,
	MR, NE, SN,			
•	AU 2005247108	A1 20051208	AU 2005-247108	20050531
	CA 2533061	AA 20051208	CA 2005-2533061	20050531
		A1 20060510		
PRIO	RITY APPLN. INFO.:		ES 2004-1312	
		•	WO 2005-EP1969	
		:	WO 2005-GB722	
AB	A combination which	comprises (a) a	WO 2005-EP5841	
AD	of M3 muscarinic re	comprises (a) a	PDE4 inhibitor and (3R)-1-phenethyl-3-	(b) an antagonist
	carbonylovy) -1-azon	iabicyclo[2 2 2]	octane, in the form of	yn-xantnene-y-
	anion X, which is a	nharmaceuticall	y acceptable anion of	or a sait having an
	polyvalent acid.	pharmaccacrearr	y deceptable anion of	. a mono or
IT	257892-33-4			
		gical activity):	THU (Therapeutic use	e) · BIOL
	(Biological study);		in (inclupeded do	2, , 2102
•			carinic agents and PI	E4 inhibitors)
RN	257892-33-4 CAPLUS		J	,
CN	1H-Indole-3-acetami	de, N-(3,5-dichl	oro-4-pyridinyl)-1-[(4 -
			o- (9CI) (CA INDEX N	

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 6 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1267950 CAPLUS

DOCUMENT NUMBER: 144:266281

TITLE: AWD-12-281 (inhaled) Elbion/GlaxoSmithKline

AUTHOR(S): Gutke, Hans-Juergen; Guse, Jan-Hinrich; Khobzaoui,

Moussa; Renukappa-Gutke, Thejavathi; Burnet, Michael

CORPORATE SOURCE: Synovo GmbH, Tubingen, D-72076, Germany

SOURCE: Current Opinion in Investigational Drugs (Thomson

Scientific) (2005), 6(11), 1149-1158

CODEN: COIDAZ; ISSN: 1472-4472

PUBLISHER: Thomson Scientific DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. Elbion (formerly ASTA Medica) and GlaxoSmithKline are developing an inhaled formulation of AWD-12-281 for the potential treatment of chronic obstructive pulmonary disease (COPD). By May 2005, phase II trials of this 5-hydroxyindole PDE4 inhibitor for COPD were ongoing.

IT **257892-33-4**, AWD-12-281

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(phosphodiestearase 4 inhibitor AWD-12-281 was safe and effective in treatment of chronic obstructive pulmonary disease patient)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

REFERENCE COUNT: 62 THERE ARE 62 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 7 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1155523 CAPLUS

DOCUMENT NUMBER: 143:416252

TITLE: Novel medicament combinations for the treatment of

respiratory diseases

PATENT ASSIGNEE(S): Boehringer Ingelheim International GmbH, Germany

SOURCE: U.S. Pat. Appl. Publ., 50 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.						DATE			APP	LICAT	ION	NO.		D	ATE	
`						-									-		
US	2005	2397	78		A1		2005	1027		US	2005-	1090	94		2	0050	419
DE	1020	0401	9540		A1		2005	1110		DE	2004-	1020	0401	9540	2	0040	422
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											2005-						
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											, EC,						
		-	-								, JP,	•	•	•	•	•	•
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	LC, LK, LI NI. NO. N				•	-	•	•				•	•	•	•	•	
	NI, NO, NZ													•		•	•
		SM,	SY,	ТJ,	TM,	TN,	TR,	TT,	TZ,	UA	, UG,	US,	UZ,	VC,	VN,	YU,	ZA,
		ZM,	zw														
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			ΝE,		•												
PRIORITY	PRIORITY APPLN. INFO.:									DE	2004-	1020	0401	9540	A 2	0040	422
										US	2004-	5785	42P		P 2	0040	610
										DE	2004-	1020	0405	2987	A 2	0041	103
											2005-					0050	
OBUED CO	STREET COLDER (C).					D 3 CT	142	41.60		L F	2005-	2270			. 2	0050.	201
OTHER SC	HER SOURCE(S):				MAR.	PAT.	143:	4162	52								

O GI

AB The present invention relates to a pharmaceutical composition comprising one or more compds. of formula I wherein n denotes 1 or 2; R1 denotes hydrogen, halogen, C1-C4-alkyl or -O-C1-C4-alkyl; R2 denotes hydrogen, halogen, C1-C4-alkyl or -O-C1-C4-alkyl; R3 denotes C1-C4-alkyl, OH, halogen, -O-C1-C4-alkyl, -O-C1-C4-alkylene-COOH, -O-C1-C4-alkylene-CO-O-C1-C4alkyl, and at least one other active substance for the treatment of respiratory diseases. The second active substance can by an anticholinergic, a phosphodiesterase IV inhibitor, a steroid, a LTD4 antagonist or an EGFR inhibitor.

257892-33-4, AWD-12-281 IT

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

I

(phosphodiesterase IV inhibitor; novel medicament combinations for treatment of respiratory diseases)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CN fluorophenyl)methyl]-5-hydroxy-α-οxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & &$$

CAPLUS COPYRIGHT 2006 ACS on STN L25 ANSWER 8 OF 40

ACCESSION NUMBER:

2005:823606 CAPLUS

DOCUMENT NUMBER:

143:206419

TITLE:

Treatment of rhinitis with anticholinergics alone or in combination with antihistamines, phosphodiesterase

4 inhibitors, or corticosteroids

INVENTOR (S):

Maus, Joachim; Petzold, Ursula; Szelenyi, Istvan;

Hoffmann, Torsten; Weingart, Mario

PATENT ASSIGNEE(S):

Sofotec G.m.b.H. & Co. K.-G., Germany PCT Int. Appl., 16 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT	NO.					DATE								D	ATE		
	WO 2005				A 2			0818					3		2	0050	124	
	WO 2005	50749	83		A 3		2006	0413				•						
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		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	
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		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	
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		RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	
		MR,	ΝE,	SN,	TD,	TG												
	US 200	52221	02		A1	:	2005	1006	1	US 2	005-	5147	0		2	0050	207	
PRIO	RITY API	PLN.	INFO	. :					1	US 2	004-	5419	50P		P 2	0040	206	
AB	The in	venti	on p	rovi	des	comb.	inat	ions	com	pris	ing	a to	pica:	l an	tich	olin	ergi	С
	drug a	lone	or i	n co	mbin	atio	n wi	th to	opic	ally	adm	inis	tere	d an	tihi	stam	ines	,

topically or orally administered phosphodiesterase 4 inhibitors or topical

corticosteroids for the treatment of rhinitis of various origins. It further comprises presentation of these combinations in locally applied formulations and includes various pharmaceutical formulations suitable for topical application, e.g. nasal sprays, nasal drops, emulsions, pastes, creams and gels.

IT **257892-33-4**, AWD-12-281

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(anticholinergics alone or in combination with antihistamines, phosphodiesterase 4 inhibitors, or corticosteroids for treatment of rhinitis)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & &$$

L25 ANSWER 9 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:823578 CAPLUS

DOCUMENT NUMBER:

143:229872

TITLE:

Preparation of aminopyri(mi)dinecarboxamide CB2 modulators for use in combination with PDE4 inhibitors for treating pain, immune, inflammatory and rheumatic

diseases

INVENTOR(S):

Green, Richard Howard; Brown, Andrew James; Connor, Helen Elizabeth; Eatherton, Andrew John; Giblin, Gerard Martin Paul; Jandu, Karamjit Singh; Knowles, Richard Graham; Mitchell, William Leonard; Naylor, Alan; O'Shaughnessy, Celestine Theresa; Palombi, Giovanni; Rawlings, Derek Anthony; Slingsby, Brian Peter; Tralau-Stewart, Catherine Jane; Whittington,

Glaxo Group Limited, UK; Doughty, Jennifer Margaret

Andrew Richard; Williamson, Richard Alexander

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 192 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO: DATE

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WO 2005074939
                          Α1
                                20050818
                                            WO 2005-GB348
                                                                    20050201
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW, ML,
             MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                            GB 2004-2355
                                                                 A 20040203
OTHER SOURCE(S):
                         MARPAT 143:229872
GΙ
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$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & &$$

The invention is related to combination of one or more CB2 modulators of AΒ formula I [X = CH, N; Y = (un)substituted Ph; R1 = H, cyclo/alkyl,(un)substituted haloalkyl; R2 = C(R7)2R3; R3 = (un)substituted non-aromatic heterocyclyl, cycloalk(en)yl, 5-6 membered aromatic heterocyclyl, etc.; R4 = H, COMe, SO2Me, cyclo/alkyl, (un) substituted haloalkyl; R6 = Me, Cl, CHmFn; n = 1-3; m = 0-2; (n + m) = 3; R7 = H, alkyl; when X = CH, R6 = Cl,or (un) substituted alkyl and R10 = H, or R10 = Cl, or (un) substituted alkyl and R10 = H; and their pharmaceutically acceptable salts] and one or more PDE4 inhibitors useful for treating conditions which are mediated by the activity of CB2 receptors or conditions which are mediated by PDE4, such as an immune disorder, an inflammatory disorder, pain, rheumatoid. The invention is also related to the preparation of CB2 modulators I. For example, reacting cyclobutylamine with 6-(2,3-dichlorophenylamino)-4trifluoromethylnicotinic acid (preparation given) gave II in 81% yield. Selected I had EC50 values of >300 nM but <1000 nM and efficacy value of >50% at the cloned human cannabinoid CB2 receptor. Three formulations are given.

IT 257892-33-4

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

II

Saloni Sharma 05/23/2006

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LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
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             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
     AU 2004241749
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                          A1
                                             AU 2004-241749
                                                                     20040519
     CA 2525946
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                                             CA 2004-2525946
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     EP 1628682
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                                             EP 2004-766017
                                                                     20040519
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             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR
     US 2006094723
                          A1
                                 20060504
                                             US 2005-556888
                                                                     20051115
     NO 2005005941
                          Α
                                 20051214
                                             NO 2005-5941
                                                                     20051214
PRIORITY APPLN. INFO.:
                                             EP 2003-11609
                                                                     20030522
                                             WO 2004-EP50869
                                                                     20040519
GΙ
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The invention relates to the combined administration of a PDE4 inhibitor and a PDE5 inhibitor for the treatment of a disease in which phosphodiesterase 4 (PDE4) and/or phosphodiesterase 5 (PDE5) activity is detrimental. Patients were administered orally one tablet of Roflumilase and once daily a tablet of Viagra. An example of another selected PDE4 inhibitor is I.

IT 257892-33-4, AWD-12-281

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (composition comprising a PDE4 inhibitor and a PDE5 inhibitor)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-οxο-(9CI) (CA INDEX NAME)

Ι

AB A pharmaceutical composition comprises: (a) a compound of formula I wherein Xis

an anion with a single neg. charge; and (b) a PDE IV inhibitor, or an enantiomer, mixture of enantiomers, racemate, solvate, or hydrate thereof. A processes for preparing them, and their use in the treatment of respiratory complaints is also disclosed. A suspension aerosol contained I bromide 0.050, AWD-12-281 0.060, soya lecithin 0.2 and TG 134a:TG 227 (2:3) q.s.

IT 257892-33-4, AWD-12-281

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (medicaments for inhalation comprising anticholinergic and PDE IV inhibitor)

·RN 257892-33-4 CAPLUS

· CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl) methyl] -5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

L25 ANSWER 14 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:1036929 CAPLUS

DOCUMENT NUMBER:

142:16825

TITLE:

Composition comprising a PDE4 inhibitor and a PDE5

inhibitor

INVENTOR(S):

Dunkern, Thorsten; Hatzelmann, Armin; Schudt,

Christian; Grimminger, Friedrich; Ghofrani, Hossein

Ardeschir

PATENT ASSIGNEE(S):

Altana Pharma A.-G., Germany

SOURCE:

PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT	NO.			KIN	D .	DATE			APPL	ICAT:	ION I	NO.		D	ATE	
WO 2004103407 WO 2004103407				A2 A3		2004 2005		ı	WO 2	004-	EP5 _. 0	869		2	0040	519
W :	W: AE, AG, AL, AM, A' CN, CO, CR, CU, C' GE, GH, GM, HR, H		CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		

(PDE4 inhibitor, combination therapy agent; preparation of aminopyri(mi)dinecarboxamide CB2 modulators for use in combination with PDE4 inhibitors for treating pain, immune, inflammatory and rheumatic diseases)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 10 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:735067 CAPLUS

DOCUMENT NUMBER:

143:186747

TITLE:

Combination of anticholinergics and inhibitors of

phosphodiesterase type 4 for the treatment of

respiratory diseases

INVENTOR(S):

Maus, Joachim; Cnota, Peter Jurgen; Szelenyi, Istvan;

Fyrnys, Beatrix

PATENT ASSIGNEE(S):

Sofotec GmbH & Co. Kg, Germany

SOURCE: U.S. Pat. Appl. Publ., 5 pp.

CODEN: USXXCO

DOCUMENT TYPE:

. Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT	PATENT NO. 				DATE			APPL	ICAT:	ION 1	NO.		D	ATE		
		-		-	~								-			
US 2005	175547		A1		2005	0811		US 2	005-	5146	3		2	00502	207	
WO 2005	074982		A2	:	2005	0818	1	WO 2	005-1	EP65	1		2	0050	124	
WO 2005	074982		A3	:	2006	0406										
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RW:	BW, GH	I, GM,	ΚE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	ŪĠ,	ZM,	ZW,	AM,	
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RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.:

US 2004-541955P The present invention relates to a combination of an inhaled/oral PDE 4 inhibitor in combination with inhaled anticholinergic bronchodilators (muscarinic receptor antagonists), preferentially roflumilast or AWD-12-281 and R,R-glycopyrrolate, for symptomatic or prophylactic treatment of respiratory diseases, especially those accompanied by obstruction or inflammation such as chronic obstructive pulmonary disease or asthma. It further comprises the presentation of this combination in a locally applied (inhaled) formulation and application in an inhalation device for instance in the Novolizer. The influence of R,R-glycopyrrolate in combination with PDE4 inhibitors on TNF secretion was investigated by using human peripheral blood mononuclear cells. Powder inhalation with 50

prepared IT 257892-33-4, Awd 12 281

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

μg R,R-glycopyrrolate and 500 μg AWD 12-281 per single dose were

(combination of anticholinergics and inhibitors of phosphodiesterase for treatment of respiratory diseases)

257892-33-4 CAPLUS RN

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CNfluorophenyl) methyl] -5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

ANSWER 11 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:586215 CAPLUS

DOCUMENT NUMBER:

143:120526

Pharmaceutical compositions based on anticholinergics TITLE:

and additional active ingredients

INVENTOR (S):

Pairet, Michel; Pieper, Michael P.; Meade, Christopher John Montague; Reichl, Richard; Schmelzer, Christel;

Jung, Birgit

PATENT ASSIGNEE(S):

Boehringer Ingelheim Pharma GmbH & Co. Kg, Germany

U.S. Pat. Appl. Publ., 50 pp., Cont.-in-part of U.S. Ser. No. 824,391.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
TG 2005140562		20050705		-	
US 2005148562 DE 10062712	A1 Al	20050707	US 2004-6940		20041208
DE 10062712 DE 10063957	A1	20020620 20020627	DE 2000-10062712 DE 2000-10063957		20001215
DE 10003337 DE 10110772	A1	20020827	DE 2000-10063937 DE 2001-10110772		20001220 20010307
DE 10110772 DE 10111058	A1	20020912	DE 2001-10110772 DE 2001-10111058		20010307
DE 10111036 DE 10113366	A1	20020926	DE 2001-10111038 DE 2001-10113366		20010308
DE 10113300 DE 10138272	A1	20020320	DE 2001-10113300 DE 2001-10138272		20010320
US 2002151541	A1	20030227	US 2001-7182		20010010
US 2002183292	A1	20021205	US 2001-86145		20011019
US 2002137764	A1	20020926	US 2001-40196		20011025
US 2002122773	A1	20020905	US 2001-27662		20011220
DE 10206505 °	A1	20030828	DE 2002-10206505		20020216
US 2002169181	A1	20021114	US 2002-92116		20020306
US 6620438	B2	20030916			•
US 2002193393	A1 ·	20021219	US 2002-93240		20020307
US 2002183347	À1	20021205	US 2002-100659		20020318
US 6608054	B2	20030819			
US 2003158196	. A1	20030821	US 2003-360064		20030207
US 2003181478	A1	20030925	US 2003-395777		20030324
US 6890517 US 2003203925	B2	20050510	110 2002 412065		00000111
US 2003203925	A1 A1	20031030 20031113	US 2003-413065 US 2003-419358		20030414
US 6696042	B2 .	20031113	05 2003-419356		20030421
US 2004024007	A1		US 2003-613783		20030703
US 2004151770	A1	20040805	US 2004-763894		20030703
US 2004161386	A1	20040819	US 2004-775901		20040123
US 2004176338	A1	20040909	US 2004-776757		20040211
US 2004192675	A1	20040930	US 2004-824391		20040414
US 2005147564	A1	20050707	US 2005-68134		20050228
PRIORITY APPLN. INFO.:			DE 2000-10054042	Α	20001031
			US 2000-253613P	P	20001128
			DE 2000-10062712	Α	20001215
		•	DE 2000-10063957	Α	20001220
			US 2000-257220P	P	20001221
			US 2000-257221P	P	20001221
•			DE 2001-10110772	A	20010307
			DE 2001-10111058	A	20010308
•			DE 2001-10113366 US 2001-281653P	A	20010320
			US 2001-281857P	P P	20010405 20010405
•			US 2001-281874P	P	20010405
			DE 2001-10138272	A	20010403
			US 2001-314599P	P	20010010
			US 2001-7182		20011019
•			US 2001-86145		20011019
·		•	US 2001-27662		20011220
			DE 2002-10206505	Α	20020216
			US 2002-92116	A1	20020306
			US 2002-93240		20020307
			US 2002-100659		20020318
			US 2002-369213P	P	20020401
		,	US 2003-360064		20030207
			US 2003-413065		20030414
			US 2003-419358		20030421
			US 2003-613783	A2	20030703

US	2004-763894	A2	20040123
US	2004-775901	A2	20040210
US	2004-776757	A2	20040211
US	2004-824391	A2	20040414
US	2001-40196	B1	20011025
US	2003-395777	`A1	20030324

OTHER SOURCE(S): MARPAT 143:120526

AB A pharmaceutical composition comprising an anticholinergic and at least one addnl. active ingredient selected from among corticosteroids, dopamine agonists, PDE-IV inhibitors, NK1-antagonists, endothelin antagonists, antihistamines, and EGFR-kinase inhibitors, processes for preparing them and their use in the treatment of respiratory diseases. Among a number of compds. prepared was N-[2-[3,5-bis(trifluoromethyl)phenyl]ethyl]-2-[4-[(3-hydroxypropyl)methylamino]piperidin-1-yl]-N-methyl-2-phenylacetamide. Inhalable powders include a formulation containing tiotropium bromide, budesonide, and lactose.

IT 257892-33-4, AWd-12-281

RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. based on anticholinergics and addnl. active ingredients)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & \\ & & & \\ & &$$

L25 ANSWER 12 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:136543 CAPLUS

DOCUMENT NUMBER: 142:246142

TITLE: Medicaments comprising PDE IV inhibitors and an

anticholinergic agent for treating respiratory

disorders

INVENTOR(S): Germeyer, Sabine; Meade, Christopher John Montague;

Meissner, Helmut; Morschhaeuser, Gerd; Pairet, Michel;

Pestel, Sabine; Pieper, Michael P.; Pohl, Gerald;

Reichl, Richard; Speck, Georg

PATENT ASSIGNEE(S): Boehringer Ingelheim International G.m.b.H., Germany;

Boehringer Ingelheim Pharma G.m.b.H. & Co. K.-G.

SOURCE: PCT Int. Appl., 41 pp.

CODEN: PIXXD2 ·

DOCUMENT TYPE: Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA	TENT 1													ATE			
WO	2005																
	W:	ΑE,	AG,	ΑL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,
							LV,	•							-		-
							PL,						-	-	-	-	•
							TZ,								-		
	RW: BW, GH,													-	-		
	AZ, BY, KG,																
							GR,										
		-			•		CF,	-		-			•			•	•
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	2533																
	1651															0040	
Li							ES,										
	к.						TR,							мп,	SE,	MC,	Ρ1,
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PRIORIT	PRIORITY APPLN. INFO.:										003-:					0030	
									1	US 2	003-	5081	19P]	P 2	0031	002
									1	WO 2	004-1	EP80	03	V	<i>N</i> 2	0040	723
OTHER S	OURCE		MAR	$\mathbf{T}\mathbf{A}\mathbf{q}$	142:2	24614	12										

The present invention relates to pharmaceutical compns. based on PDE ${\tt IV}$ AB inhibitors and salts of a novel anticholinergic, processes for preparing them and their use in the treatment of respiratory complaints. For example, scopine 9-methylfluorene-9-carboxylate methobromide was prepared and formulated into inhalable powder containing the drug 80 μ g, AWD-12-281 200 $\mu g,$ and lactose 12220 μg per capsule.

IT 257892-33-4, AWD 12-281

> RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (GW-842470; inhalable compns. comprising anticholinergic agent and PDE IV inhibitors for treating respiratory disorders)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CNfluorophenyl) methyl] -5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 13 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2005:99152 CAPLUS 142:204737

DOCUMENT NUMBER: TITLE:

Medicaments for inhalation comprising an anticholinergic and a PDE IV inhibitor

INVENTOR(S):

Meade, Christopher John Montague; Pairet, Michel;

Pieper, Michel; Pieper, Michael P.

PATENT ASSIGNEE(S):

Boehringer Ingelheim International G.m.b.H., Germany

SOURCE:

U.S. Pat. Appl. Publ., 18 pp.

CODEN: USXXCO

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PA'	TENT 1	NO.	•		KIND DATE				APPLICATION NO.						DATE				
US	2005	0268	86		A1		2005	0203	,	US 2	004-	8915	51		20	040	715		
CA	2534	125			·AA 20050217					CA 2	004-	2534	20040717						
WO	WO 2005013993					A1 20050217			1	WO 2	004-1	EP80:	20040717						
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,		
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	.KR,	ΚZ,	LC,		
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,		
		NO,	ΝZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,		
		TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW		
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	ŪĠ,	ZM,	ZW,	AM,		
							RU,												
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,		
		SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	ΝE,		
		SN,	TD,	TG															
EP	1651	222			A1		2006	0503		EP 2	004-	7411	28		2 (040	717		
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
		ΙE,	SI,	FI,	RO,	CY,	TR,	BG,	CZ,	EE,	HU,	PL,	SK						
PRIORIT	Y APP	LN.	INFO	. :						EP 2	003-	1716	4	1	A 20	0030	729 ⁻		
						US 2003-508125P]	P 20	0031	002					
									,	WO 2	004-1	EP80	24	7	W -20	0040	717		
OTHER S		MAR	PAT	142:	2047	37													

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$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

L25 ANSWER 15 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

2004:996001 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 141:406065

TITLE: Composition comprising a PDE-4 inhibitor and a

TNF-alpha antagonist

INVENTOR(S): Barsig, Johannes; Weimar, Christian

PATENT ASSIGNEE(S): Altana Pharma AG, Germany SOURCE: PCT Int. Appl., 29 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

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PATENT NO.
                       KIND
                               DATE
                                             APPLICATION NO.
                                                                         DATE
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WO 2004098633
                       A1
                               20041118
                                            WO 2004-EP50748
                                                                        20040510
    W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
         CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
         GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
         LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
         NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
         \mathtt{TJ},\ \mathtt{TM},\ \mathtt{TN},\ \mathtt{TR},\ \mathtt{TT},\ \mathtt{TZ},\ \mathtt{UA},\ \mathtt{UG},\ \mathtt{US},\ \mathtt{UZ},\ \mathtt{VC},\ \mathtt{VN},\ \mathtt{YU},\ \mathtt{ZA},\ \mathtt{ZM},\ \mathtt{ZW}
    RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
         AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
         EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
         SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
         SN, TD, TG
                                              EP 2003-10581
                                                                     A 20030512
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PRIORITY APPLN. INFO.:

The invention relates to the combined administration of a PDE4 inhibitor and a $TNF\alpha$ antagonist selected from the group consisting of etanercept, onercept and pegsunercept for the treatment of a disease in which phosphodiesterase 4 (PDE4) and/or tumor necrosis factor alpha $(TNF\alpha)$ activity is detrimental.

IT 257892-33-4, AWD 12-281

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(therapeutic activity of phosphodiesterase 4 inhibitors and $TNF\alpha$ antagonists)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CN

fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ &$$

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 16 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

4

ACCESSION NUMBER:

2004:995979 CAPLUS

DOCUMENT NUMBER:

141:406064

TITLE:

Composition comprising a PDE4 inhibitor and soluble human Type II interleukin-1 receptor (shuIL-1RII) for

disease therapy

INVENTOR (S):

Barsig, Johannes

PATENT ASSIGNEE(S):

Altana Pharma AG, Germany

SOURCE: PCT Int. Appl., 24 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT	NO.	٠.,		KIN		DATE		1	APPL:	ICAT:	ION 1	NO.		D	ATE	·
	WO 200	40986	06				2004	 1118	1	WO 2	 004-1	EP50'	749		2	0040	 510
	W:		AG,														
			co,														
			GH,														
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			NZ,														
			TM,														
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			BY,														
			ES,														
			SK,		BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	·ML,	MR,	NE,
			TD,														
PRIO	RITY AP	PLN.	INFO	. :					1	EP 20	003-:	1059	5		A 2	0030	512
AB	The in	venti	on r	elat	es t	o th	e co	mbin	ed a	dmin:	istra	atio	n of	a P	DE4	inhil	bitor
	and sh	uIL-1	R II	for	the	tre	atme:	nt o	fa (disea	ase :	in wl	hich	pho	spho	dies	terase
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IT	257892										•						
	RL: PA	C (Ph	arma	colo	qica	l ac	tivi	tv);	THU	(The	erap	euti	c use	e):]	віоь		
	(Biolo				_									, , .			

(composition comprising a PDE4 inhibitor and soluble human Type II interleukin-1 receptor (shuIL-1RII) for disease therapy)

RN257892-33-4 CAPLUS

CN

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: \ RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2006 ACS on STN L25 ANSWER 17 OF 40

ACCESSION NUMBER:

2004:995978 CAPLUS

DOCUMENT NUMBER:

141:406063

TITLE:

Pharmaceutical composition comprising a PDE4 inhibitor

and IL-1 trap for treatment of disease

INVENTOR(S):

Barsig, Johannes

PATENT ASSIGNEE(S):

Altana Pharma AG, Germany

SOURCE:

PCT Int. Appl., 24 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	ENT 1	NO.			KIN	D 1	DATE			APPL	ICAT	ION	NO.		D	ATE		
WO	2004	0986	05		A1		2004	1118	WO 2004-EP50747						20040510			
	W :	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,	
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	.LC,	
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,	
•		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	
		TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	ŪĠ,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW	
	RW:	BW,	GH,	GM,	KΕ,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	
		ΑZ,	BY,	KG,	KZ,	MD,	RU,	ТJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	
		SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	
		SN,	TD,	TG														

PRIORITY APPLN. INFO.:

EP 2003-10631 A 20030512

The invention relates to the combined administration of a PDE4 inhibitor and IL-1 Trap for the treatment of a disease in which phosphodiesterase 4 (PDE4) and/or interleukin-1 (IL-1) activity is detrimental.

257892-33-4, AWD 12-281 IT

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(pharmaceutical composition comprising a PDE4 inhibitor and IL-1 trap for treatment of disease)

RN257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CN fluorophenyl) methyl] -5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 18 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:610086 CAPLUS

DOCUMENT NUMBER:

141:134069

TITLE:

PDE4 inhibitors for the treatment of neoplasms of

lymphoid cells

INVENTOR (S):

Hatzelmann, Armin; Tenor, Hermann; Gekeler, Volker;

Sanders, Karl; Garattini, Enrico; Braunger, Juergen;

Schudt, Christian

PATENT ASSIGNEE(S):

Altana Pharma Ag, Germany

SOURCE:

PCT Int. Appl., 78 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 2004062671	A2 20040729	WO 2004-EP196	20040114
WO 2004062671	A3 20050127		
W: AE, AG, A	L, AM, AT, AU, AZ,	BA, BB, BG, BR, BW, BY,	BZ, CA, CH,
CN, CO, C	R, CU, CZ, DE, DK,	DM, DZ, EC, EE, EG, ES,	FI, GB, GD,
GE, GH, G	M, HR, HU, ID, IL,	IN, IS, JP, KE, KG, KP,	KR, KZ, LC,
LK, LR, L	S, LT, LU, LV, MA,	MD, MG, MK, MN, MW, MX,	MZ
AU 2004204355	A1 20040729	AU 2004-204355	20040114
CA 2512819	AA 20040729	CA 2004-2512819	2.0040114
EP 1587512	A2 20051026	EP 2004-701902	20040114
R: AT, BE, C	H, DE, DK, ES, FR,	GB, GR, IT, LI, LU, NL,	SE, MC, PT,
IE, SI, L	T, LV, FI, RO, MK,	CY, AL, TR, BG, CZ, EE,	HU, SK

PRIORITY APPLN. INFO.:

EP 2003-787 WO 2004-EP196 20030114 20040114

OTHER SOURCE(S):

MARPAT 141:134069

The invention relates to the use of certain PDE4 inhibitors alone or in combination with one or more differentiation inducing agents and/or an agent effective in raising intracellular concns. of cAMP or a stable analog of cAMP in the preparation of pharmaceutical compns. for the treatment of neoplasms of lymphoid cells.

257892-33-4, AWD-12-281 IT

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(phosphodiesterase 4 (PDE4) inhibitors for treatment of neoplasms of lymphoid cells in combination with differentiation inducers and agents that increase cAMP levels or cAMP analogs)

RN257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CN fluorophenyl) methyl] -5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} F \\ \hline \\ CH_2 \\ \hline \\ N \\ \hline \\ C-C-NH \\ \hline \\ C1 \\ \end{array}$$

L25 ANSWER 19 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:467725 CAPLUS

DOCUMENT NUMBER:

141:17651

TITLE:

Phosphodiesterase IV and phosphodiesterase III/IV inhibitors for use in the treatment of cachexia

INVENTOR (S):

Schmidt, Mathias

PATENT ASSIGNEE(S):

Altana Pharma A.-G., Germany

SOURCE:

PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PAT	CENT 1	NO.		•	KIN	D	DATE		APPLICATION NO.						DATE		
						-		- 							_		
WO	2004	0478	17		A1		2004	0610	WO 2003-EP13313						20031126		
	W:	ΑE,	ΑL,	ΑU,	BA,	BR,	CA,	CN,	CO,	DZ,	EC,	EG,	GE,	HR,	ID,	IL,	IN,
							MA,										
		VN,	YU,	ZA,	ZW												
	RW:	AM,	AZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,
							GB,										

SI, SK, TR CA 2506949 AA 20040610 CA 2003-2506949 20031126 AU 2003289898 A1 20040618 AU 2003-289898 20031126 EP 1567136 A1 20050831 EP 2003-782232 20031126 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK T2 20060316 JP 2004-554493 JP 2006508996 20031126 US 2005-535815 US 2006079540 **A1** 20060413 20050520 EP 2002-26548 PRIORITY APPLN. INFO.: 20021127 W WO 2003-EP13313 20031126

The invention discloses the use of a PDE IV or PDE III/IV inhibitor for AB the treatment of cachexia.

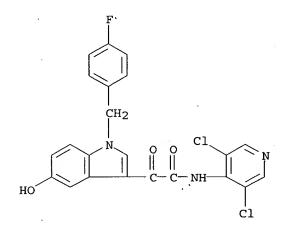
IT 257892-33-4

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(phosphodiesterase IV and phosphodiesterase III/IV inhibitors for treatment of cachexia)

RN257892-33-4 CAPLUS

CN1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 20 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:203704 CAPLUS

DOCUMENT NUMBER:

140:229455

TITLE:

Combination of glucocorticoids and PDE-4-inhibitors

for treating respiratory diseases, allergic diseases,

asthma and COPD

INVENTOR (S):

Locher, Mathias; Hermann, Robert

PATENT ASSIGNEE(S):

Viatris G.m.b.H. & Co. K.-G., Germany

SOURCE:

PCT Int. Appl., 26 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent German

LANGUAGE: FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND · DATE APPLICATION NO. DATE

```
WO 2004019984
                          A1
                                20040311
                                            WO 2003-EP8607
         W: AU, BR, CA, CN, CO, CZ, GE, HR, ID, IL, IN, JP, KR, LT, LV, MD,
             MK, MX, NO, NZ, PL, SG, UA, US, UZ, YU, ZA
         RW: AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE,
             DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE,
             SI, SK, TR
    CA 2492645
                          ΔΔ
                                20040311
                                            CA 2003-2492645
                                                                    20030804
    AU 2003255365 ,
                                20040319
                          A1
                                            AU 2003-255365
                                                                    20030804
     EP 1526870
                                20050504
                                            EP 2003-790851
                          A1
                                                                    20030804
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, TR, BG, CZ, EE, HU, SK
     JP 2005539042
                          T2
                                20051222
                                            JP 2004-531853
                                                                    20030804
     US 2005288265
                          A1
                                20051229
                                            US 2005-523802
                                                                    20050209
    NO 2005001212
                          Α
                                20050308
                                            NO 2005-1212
                                                                    20050308
PRIORITY APPLN. INFO.:
                                            DE 2002-10236688
                                                                 Α
                                                                    20020809
                                            WO 2003-EP8607
                                                                 W
                                                                   20030804
AΒ
     The invention relates to a novel combination of a glucocorticoid, especially
     loteprednol, and at least one phosphodiesterase-4 inhibitor
     (PDE-4-inhibitor), especially hydroxyindole-derivative
N-(3,5-dichloropyridine-4-yl)-
     2-[1-(4-fluorbenzyl)-5-hydroxyindole-3-yl]-2-oxoacetamide, for a
     simultaneous, sequential or sep. administration in the treatment of
    respiratory diseases, allergic diseases, asthma and chronic obstructive
     pulmonary diseases (COPD). Formulation of glucocorticoids and
     PDE-4-inhibitors can be prepared sep. and applied at the same time or at
     different times during the day; also combinations can be formulated.
TT
     257892-33-4
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (combination of glucocorticoids and PDE-4-inhibitors for treating
        respiratory diseases, allergic diseases, asthma and COPD)
RN
     257892-33-4 CAPLUS
```

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 21 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:

2004:120846 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-

fluorophenyl) methyl] -5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

DOCUMENT NUMBER:

140:163707

CN

TITLE:

Method for producing highly pure hydroxyindolylglyoxylic acid amides

INVENTOR(S):

Jaensch, Hans-Joachim; Hartenhauer, Helge; Stange,

Hans; Hoefgen, Norbert; Schaefer, Juergen

PATENT ASSIGNEE(S):

Elbion AG, Germany PCT Int. Appl., 32 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATE	ENT NO	Э.			KIND DATE			7	APPL	ICAT		DATE					
						-					-				_		
	200401																
	W: 7	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	· IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
	I	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NI,	NO,	NZ,	OM,
											YU,						
	RW: C	ЗН,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
											NL,						
	I	BF,	вJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG
CA 2	249398										2003-						
AU 2	200325	5534	1		A1		2004	0223		AU 2	2003-	2553	41		2	0030	731
BR 2		•															
	16716																
JP 2	20065	0300	2		Т2												
																0050	
																0020	801
											2003-						
OTHER SOL	THER SOURCE(S):						T 14	0:16						707			
GI	(- , -											•				
CA 2 AU 2 US 2 EP 1 BR 2 CN 1 JP 2 NO 2 PRIORITY	RW: 0 1 249398 200329 200400 152519 R: 2 200300 167169 200650 200500 APPLI	PG, TR, GH, KG, FI, BF, 82 5534 6393 97 AT, 1346 92 0300 0108 N. I	PH, TT, GM, KZ, FR, BJ, 1 9 BE, SI, 67	PL, TZ, KE, MD, GB, CF,	PT, UA, LS, RU, GR, CG, AA A1 A1 DE, LV, A T2 A	RO, UG, MW, TJ, HU, CI,	RU, US, MZ, TM, IE, CM, 2004 2005 ES, RO, 2005 2005 2006 2005	SC, UZ, SD, AT, IT, GA, 0212 0223 0401 0427 FR, MK, 0705 0921 0126 0502	SD, VC, SL, BE, LU, GN,	SE, VN, SZ, BG, MC, GQ, CA 2 EP 2, AL, BR 2 CN 2 EP 2 NO 2 SWO 2	SG, YU, TZ, CH, NL, GW, 2003-2003- 1T, TR, 2003- 2004- 2005- 2002- 2003-	SK, ZA, UG, PT, ML, 2493 2553 6314 7663 LI, BG, 1346 8185 5254 1086 4002 EP85	SL, ZM, ZM, CZ, RO, 982 41 75 89 LU, CZ, 7 37 09	SY, ZW, DE, SE, NE,	TJ, AM, DK, SI, SN, 2 2 2 SE, HU, 2 2 2 P 2	TM, AZ, EE, SK, TD, 0030 0030 MC, SK 0030 0030 0030 0030	TN BY ES TR TG 731 731 731 731 731 228 801

$$\begin{array}{c|c} R^2 & & & O \\ \hline & & & & \\ R^3 & & & & \\ R_1 & & & & \\ \end{array}$$

I

The invention relates to a method for producing hydroxyindolylglyoxylic acid amides I [R1 = (un)branched, (un)saturated C1-6-alkyl, 3- to 14-membered mono-, bi- or tricarbocycle, (un)substituted 5- to 15-membered heterocycle (1 - 6 heteroatoms - N, O, S); R2, R3 = H, OH (one or both OH); R4 = (un)substituted mono- or polycyclic aromatic C6-14-carbocycle, 5 to 15-membered heterocycle (containing N, O, S)] in high yields and in a particularly pure form from 5- or 6-benzyloxyindole or

5,6-di(benzyloxy) indole compds. The method comprises: (a) reaction of 5-or 6-benzyloxyindole or 5,6-di(benzyloxy) indole with R1X (X = halogen); (b) C-acylation of the 1-substituted indole with (COX)2; (c) reaction of the [indol-3-yl]glyoxyl halide with NH3, NH2R4, NH(R4)2; and (d) hydrogenolytic debenzylation. Thus, AWD 12-281 [I; R1 = CH2C6H4F-4, R2 = OH, R3 = H, R4 = 3,5-dichloro-4-pyridyl] was prepared from 5-(benzyloxy) indole via N-benzylation with 4-FC6H4CH2Cl, C-acylation with (COCl)2, amidation with 4-amino-3,5-dichloropyridine, and hydrogenolytic debenzylation of I [R1 = CH2C6H4F-4, R2 = OCH2Ph, R3 = H, R4 = 3,5-dichloro-4-pyridyl].

IT 247584-23-2P 247584-24-3P 247584-26-5P 247584-27-6P 247584-28-7P 247584-29-8P 247584-30-1P 247584-31-2P 247584-32-3P 257892-33-4P, AWD 12-281 656237-82-0P RL: PUR (Purification or recovery); SPN (Synthetic preparation); PREP

(Preparation)
(preparation of highly pure hydroxyindolylglyoxylic acid amides)
247584-23-2 CAPLUS

RN 247584-23-2 CAPLUS
CN 1H-Indole-3-acetamide, 1-[(2,6-difluorophenyl)methyl]-5-hydroxy-αoxo-N-4-pyridinyl- (9CI) (CA INDEX NAME)

RN 247584-24-3 CAPLUS
CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(2,6-difluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

RN 247584-26-5 CAPLUS
CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-hydroxy-α-oxo1-propyl- (9CI) (CA INDEX NAME)

RN 247584-27-6 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-hydroxy-1-(1-methylethyl)-α-oxo- (9CI) (CA INDEX NAME)

RN 247584-28-7 CAPLUS

CN 1H-Indole-3-acetamide, 1-(cyclopentylmethyl)-N-(3,5-dichloro-4-pyridinyl)-5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

$$C1$$
 NH
 $C=0$
 $C=0$
 $C=0$

RN 247584-29-8 CAPLUS

CN 1H-Indole-3-acetamide, N-(2,6-dichlorophenyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

RN 247584-30-1 CAPLUS

CN 1H-Indole-3-acetamide, N-[2,6-dichloro-4-(trifluoromethyl)phenyl]-1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo- (9CI) (CÁ INDEX NAME)

RN 247584-31-2 CAPLUS

CN 1H-Indole-3-acetamide, N-[2,6-dichloro-4-(trifluoromethoxy)phenyl]-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-οxο-(9CI) (CA INDEX NAME)

HO

$$CH_2$$
 CH_2
 $C-C-NH$
 $C-C-NH$
 $C1$

RN 247584-32-3 CAPLUS

CN lH-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-6-hydroxy- α -oxo-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} F \\ \hline \\ CH_2 \\ \hline \\ N \\ C \\ C \\ C \\ N \\ \end{array}$$

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

RN 656237-82-0 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-5-hydroxy-1-[(4-methoxyphenyl)methyl]- α -oxo- (9CI) (CA INDEX NAME)

L25 ANSWER 22 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:41257 CAPLUS

TITLE:

140:87709

DOCUMENT NUMBER:

Pharmaceutical compositions comprising anticholinergic

agents and phosphodiesterase IV (PDE-IV) inhibitors

for the treatment of respiratory diseases

Pairet, Michel; Meade, Christopher John Montague; INVENTOR(S):

Pieper, Michael P.

PATENT ASSIGNEE(S):

Boehringer Ingelheim Pharma G.m.b.H. & Co. K.-G.,

Germany

SOURCE: .

PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent German '

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT N	o.	KIND	DATE	APPLICATION NO.	DATE			
WO 20040	04704	A1	20040115	WO 2003-EP6668	20030625			
				BA, BB, BG, BR, BY,				
•	CO, CR, C	U, CZ, D	DE, DK, DM,	DZ, EC, EE, ES, FI,	GB, ĠD, GE, GH,			
	GM, HR, F	U, ID, I	IL, IN, IS,	JP, KE, KG, KP, KR,	KZ, LC, LK, LR,			
				MK, MN, MW, MX, MZ,				
				SD, SE, SG, SK, SL,	TJ, TM, TN, TR,			
				VN, YU, ZA, ZM, ZW				
				SL, SZ, TZ, UG, ZM,				
				BE, BG, CH, CY, CZ,				
				LU, MC, NL, PT, RO,				
	BF, BJ, (F, CG, C	CI, CM, GA,	GN, GQ, GW, ML, MR,	NE, SN, TD, TG			
DE 10230	769	-A1	20040122	DE 2002-10230769	20020709			
				CA 2003-2492026				
				AU.2003-242755				
	76			EP 2003-762509				
R:	AT, BE, (H, DE, D	OK, ES, FR,	GB, GR, IT, LI, LU,	NL, SE, MC, PT,			
	IE, SI, I	T, LV, F	FI, RO, MK,	CY, AL, TR, BG, CZ,	EE, HU, SK			
	32379			JP 2004-518566				
US 20040	58950	A1	20040325	US 2003-614365	20030707			
PRIORITY APPL	N. INFO.			DE 2002-10230769	A 20020709			
				US 2002-407895P	P 20020903			
				WO 2003-EP6668	W 20030625			

MARPAT 140:87709 OTHER SOURCE(S):

The invention provides pharmaceutical compns. comprising anticholinergic agents and PDE-IV inhibitors, as well as a method for the production and use thereof in the treatment of respiratory diseases. Powder inhalant formulations are included.

257892-33-4, AWD-12-281 645337-16-2 IT

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. comprising anticholinergic agents and phosphodiesterase IV inhibitors for treatment of respiratory diseases)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CN fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ &$$

645337-16-2 CAPLUS RN

3-0xa-9-azoniatricyclo[3.3.1.02,4] nonane, 9,9-dimethyl-7-(1-oxo-2,2-CN diphenylpropoxy) -, bromide, $(1\alpha, 2\beta, 4\beta, 5\alpha, 7\beta)$ -, mixt. with N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5hydroxy- α -oxo-1H-indole-3-acetamide (9CI) (CA INDEX NAME)

, CM 1

> CRN 412046-80-1 CMF C24 H28 N O3 . Br

Relative stereochemistry.

Br-

CM 2

CRN 257892-33-4

CMF C22 H14 Cl2 F N3 O3

REFERENCE COUNT:

THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 23 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

9

ACCESSION NUMBER:

2003:719308 CAPLUS

DOCUMENT NUMBER:

139:240373

TITLE:

Pharmaceutical composition of a phosphodiesterase 4 (PDE4) inhibitor or a PDE3/4 inhibitor and a histamine receptor antagonist for the treatment of respiratory

diseases

INVENTOR(S):

Beume, Rolf; Bundschuh, Daniela; Weimar, Christian;

Wollin, Stefan-lutz

PATENT ASSIGNEE(S):

Altana Pharma Ag, Germany

SOURCE:

PCT Int. Appl., 87 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	ENT 1		KINI)	DATE		i.	APPL:	ICAT:	ON 1	10.		DA	ATE			
WO	20030	7405	55		A 1		2003	0912	ī	VO 20	003-I	EP187	76		20	00302	225
	W:	ΑE,	AL,	AU,	BA,	BR,	CA,	CN,	CO,	CU,	DZ,	EC,	GE,	HR,	ID,	IL,	IN,
		IS,	JP,	KR,	LT,	LV,	MA,	MK,	MX,	NO,	NZ,	PH,	PL,	SG,	TN,	UA,	US,
		VN,	YU,	ZA,	ZW												
	RW:	AM,	ΑZ,	BY,	KG,	KZ,	MD,	RU,	TJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,
		DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	SI,
		SK,	TR														
CA	24786	512			AA		2003	0912	(CA 20	003-2	24786	512		20	00302	225
	20032																
EP	1482938				A1		2004	1208]	EP 20	003-1	70813	30		20	00302	225
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		ΙE,	SI,	LT,	LV,	FI,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	SK		
BR 2003008220				Α		2005	0104]	BR 20	003-8	3220			20	00302	225	

US 2005112069 **A1** 20050526 US 2003-506875 20030225 T2 20050818 JP 2005524666 JP 2003-572572 20030225 NO 2004004230 Α 20041206 NO 2004-4230 20041006 PRIORITY APPLN. INFO.: EP 2002-4987 20020306 Α WO 2003-EP1876 W 20030225

AB The invention discloses the combined administration of PDE4 or PDE3/4 inhibitors and histamine receptor antagonists for the treatment of respiratory diseases.

IT 257892-33-4, AWD 12-281

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(phosphodiesterase 4 (PDE4) inhibitor or PDE3/4 inhibitor combination with histamine receptor antagonist for treatment of respiratory disease)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 24 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:376393 CAPLUS

DOCUMENT NUMBER:

138:379220

TITLE:

Combination of type 4 phosphodiesterase inhibitor and disease-modifying anti-rheumatic drug for treating

where the descripting and the mediant drug I

rheumatoid arthritis

INVENTOR (S):

Barsiq, Johannes

PATENT ASSIGNEE(S):

Germany

SOURCE:

U.S. Pat. Appl. Publ., 13 pp.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2003092706	A1	20030515	US 2002-184068	20020628
CA 2399840	AA	20030509	CA 2002-2399840	20020827
WO 2003039552	A1	20030515	WO 2002-EP12415	20021107

W: AE, AL, BA, BR, CN, CO, CU, DZ, EC, GE, HR, HU, ID, IL, IN, IS, JP, KR, LT, LV, MA, MK, MX, NO, NZ, PH, PL, RO, SG, SI, TN, UA, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT,

LU, MC, NL, PT, SE, SK, TR

T2

EP 1448202 Al 20040825 EP 2002-792742 20021107 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK

20050407

JP 2005508983
PRIORITY APPLN. INFO.:

JP 2003-541843 20021107 EP 2001-607 A 20011109 WO 2002-EP12415 W 20021107

AB The invention relates to the combined administration of a PDE4 or PDE3/4 inhibitor and a disease modifying anti-rheumatic drug (DMARDs) or anti-rheumatic or anti-arthritic drug. Oral treatments with Roflumilast plus methotrexate or Pumafentrine HCl plus methotrexate had additive beneficial effects in delaying the onset and reducing the severity of collagen-induced arthritis in DBA/1 mice.

IT 257892-33-4, AWD-12-281

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PDE4 or PDE3/4 inhibitor; combination of phosphodiesterase 4 inhibitor and disease-modifying anti-rheumatic drug for treating rheumatoid arthritis)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-οxο-(9CI) (CA INDEX NAME)

L25 ANSWER 25 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:242192 CAPLUS

DOCUMENT NUMBER:

138:248511

TITLE:

Combination of phosphodiesterase 4 inhibitor and nonsteroidal antiinflammatory drug in treatment of

inflammation

INVENTOR (S):

Hatzelmann, Armin; Eltze, Manfrid; Klein, Thomas;

Kley, Hans-Peter

PATENT ASSIGNEE(S):

Altana Pharma A.-G., Germany

SOURCE:

PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

PA	TENT :	NO.					ATE		- 1	APP	LICAT	ION I	NO.		· D .	ATE	
WO	2003	0244	 89		A2		003	0327	1	WO :	2002-1	EP10	 424		2	0020	 917
WO	2003	0244	89		A 3	2	003	0918									
•	W:	ΑE,	AL,	AU,	BA,	BR,	CA,	CN,	CO,	CU	, DZ,	EC,	GE,	HR,	HU,	ID,	IL,
		IN,	IS,	JP,	KR,	LT,	LV,	MA,	MK,	MX	, NO,	NZ,	PH,	PL,	RO,	SG,	SI,
		TN,	UA,	US,	VN,	ΥU,	ZA,	ZW									•
	RW:	AM,	AZ,	BY,	KG,	KZ,	MD,	RU,	TJ,	TM	, AT,	BE,	BG,	CH,	CY,	CZ,	DE,
											, LU,						-
CA	CA 2459757				AA	2	003	0327		CA :	2002-:	2459	757		2	0020	917
EP	EP 1429807				A2	2	004	0623		EP :	2002-	7723	13		2	0020	917
	EP 1429807 R: AT, BE, CH																
		ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL	, TR,	BG,	CZ,	EE,	SK		
BR	2002	0126	06		Α	2	004	0817]	BR :	2002-	1260	6		2	0020	917
JP	2005	5040	77		T2	2	005	0210		JP :	2003-	5285	83		2	0020	917
CN	1625	411			Α	2	005	0608		CN :	2002-	8182	41		2	0020	917
US	2004	2425	97		A 1	2	004	1202	1	US :	2004 -	4899	20		2	0040	318
ZA	2004	0026	54		A	2	005	0214			2004-:				_	0040	405
NO	NO 2004001596				Α	2	004	0618]	NO :	2004-	1596	•		2	0040	419
PRIORIT	RIORITY APPLN. INFO.:								:	EP :	2001-	473		7	A 2	0010	919
	IORITY APPLM. INFO.:								1	WO :	2002-1	EP10	424	1	₩ 2	0020	917

AB The invention relates to the combined administration of PDE4-inhibitors and NSAIDs for the treatment of an inflammatory disease and/or an inflammation associated disorder while minimizing gastrointestinal side effects, such as gastric erosions and ulcer, which are frequently associated with the use of NSAIDs. PDE4 inhibitors Rolipram, Roflumilast, and RP73401 inhibited or prevented diclofenac induced gastrointestinal bleeding in mice.

IT 257892-33-4, AWD 12-281

RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);
THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(phosphodiesterase inhibitor; combination of phosphodiesterase 4
inhibitor and nonsteroidal antiinflammatory drug in treatment of
inflammation)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

CAPLUS COPYRIGHT 2006 ACS on STN L25 ANSWER 26 OF 40 2003:242191 CAPLUS ACCESSION NUMBER: 138:248522 DOCUMENT NUMBER: Combined administration of phosphodiesterase PDE4 or TITLE: PDE3/4 inhibitors and leukotriene receptor antagonists for the treatment of respiratory tract disorders Beume, Rolf; Bundschuh, Daniela; Weimar, Christian; INVENTOR(S): Wollin, Stefan-Lutz Altana Pharma A .. - G., Germany PATENT ASSIGNEE(S): SOURCE: PCT Int. Appl., 40 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE _____ _ _ _ _ _____ _____ _ _ _ _ _ _ WO 2003024488 . A2 20030327 WO 2002-EP10423 20020917 WO 2003024488 **A3** 20030904 W: AE, AL, AU, BA, BR, CA, CN, CO, CU, DZ, EC, GE, HR, HU, ID, IL, IN, IS, JP, KR, LT, LV, MA, MK, MX, NO, NZ, PH, PL, RO, SG, SI, TN, UA, US, VN, YU, ZA, ZW RW: AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR CA 2460442 AA 20030327 CA 2002-2460442 20020917 EP 2002-798730 EP 1429843 Α2 .20040623 20020917 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK BR 2002012582 Α 20041013 BR 2002-12582 20020917 JP 2005505570 T2 20050224 JP 2003-528582 20020917 CN 1655846 Α 20050817 CN 2002-818260 20020917 ZA 2004002653 Α 20050214 ZA 2004-2653 NO 2004001595 Α 20040616 NO 2004-1595 US 2005014762 Α1 20050120 US 2004-489903 20040818 PRIORITY APPLN. INFO .:: EP 2001-474 20010919 WO 2002-EP10423 W 20020917 AΒ The invention relates to the combined administration of PDE4 or PDE3/4 inhibitors and leukotriene receptor antagonists for the treatment of respiratory tract disorders. The inhibitory effects of Roflumilast and Montelukast sodium salt on SRS-A-induced bronchoconstriction were additive in guinea pigs. TT 257892-33-4 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); . THU (Therapeutic use); BIOL (Biological study); USES (Uses) (phosphodiesterase inhibitor; combined administration of phosphodiesterase PDE4 or PDE3/4 inhibitors and leukotriene receptor antagonists for treatment of respiratory tract disorders)

Saloni Sharma 05/23/2006

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo-(9CI) (CA INDEX NAME)

257892-33-4 CAPLUS

RN CN

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

L25 ANSWER 27 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2003:5806 CAPLUS

DOCUMENT NUMBER:

138:78456

TITLE:

Composition comprising a PDE-4 inhibitor and

H1-receptor antagonist for treatment of respiratory

diseases

INVENTOR (S):

Knowles, Richard Graham; Ward, Peter; Nials, Anthony

Terence

PATENT ASSIGNEE(S):

SOURCE:

Glaxo Group Limited, UK

PCT Int. Appl., 18 pp.

CODEN: PIXXD2

DOCUMENT TYPE: .
LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT	NO.		KINI)]	DATE		ì	APPL	ICAT:	ION 1	10.		D/	ATE	
WO 200:	3000289		A1	•	2003	0103	1	WO 2	002-0	3B26'	79		20	00206	517
W:	AE, AG,									-		•		•	
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	LS, LT,										-				
	PL, PT,	-	-	-			-	•	SL,	TJ,	TM,	TN,	TR,	TT,	TZ,
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RW	: GH, GM,														
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CA 245	0758		AA		2003	0103	(CA 2	002-:	2450	758		20	0020	517
EP 140	1369		A1		2004	0407]	EP 2	002-	7356:	11		20	020	517
R:	AT, BE,	CH,	DE,	DK,	ES,	FR,	GB,	GŔ,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
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CN 151	3460		Α		2004	0804	(CN 2	002-	8124	73		20	00206	517
BR 200	BR 2002010473					0810]	BR 2	002-	1047	3		20	0020	517
JP 200	JP 2005501023					0113		JP 2	003-	50693	32		20	0206	517
US 200	US 2004176419					0909	1	JS 2	003-4	4809	59		20	00312	208
ZA 200	ZA 2003009587					0117		ZA 2	003-	9587			20	00312	210
PRIORITY AP	RIORITY APPLN. INFO.:						(GB 2	001-	1518:	1	7	A 20	0010	520
							1	WO 2	002-0	GB26'	79	Ţ	V 20	0020	517

AB A method of prophylaxis, treating, or reducing the duration or frequency of the exacerbations associated with a respiratory disease, such as chronic

obstructive pulmonary disease or asthma, comprises administering to a patient an effective amount of a phosphodiesterase-4 (PDE-4) inhibitor, e.g., cilomilastat, in combination with an H1-receptor antagonist, e.g., loratadine. For example, a metered dose inhaler (e.g., for 120 actuations) was prepared containing cilomilast 18 mg, loratadine 12 mg, and 1,1,1,2-tetrafluoroethane to 75.0 mg.

IT 257892-33-4

CN

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (compns. comprising PDE-4 inhibitor and H1-receptor antagonist for treatment of respiratory diseases)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-oxo- (9CI) (CA INDEX NAME)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 28 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2002:965129 CAPLUS

DOCUMENT NUMBER:

138:44711

TITLE:

Pharmaceutical compositions based on anticholinergics

and PDE-IV inhibitors

INVENTOR(S):

Pairet, Michel; Meade, Christopher J. M.; Pieper,

Michael P.

PATENT ASSIGNEE(S):

Germany

SOURCE:

U.S. Pat. Appl. Publ., 14 pp., Cont.-in-part of U.S.

Provisional Ser. No. 281,857.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

14

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 2002193393	A1	20021219	US 2002-93240		20020307
DE 10110772	A1	20020912	DE 2001-10110772		20010307
US 2004024007 ·	A1	20040205	US 2003-613783		20030703
US 2005148562	A1	20050707	US 2004-6940		20041208
PRIORITY APPLN. INFO.:			DE 2001-10110772	Α	20010307
		•	US 2001-281857P	P	20010405

DE 2000-10054042 20001031 US 2000-253613P P 20001128 DE 2000-10062712 Α 20001215 DE 2000-10063957 20001220 20001221 US 2000-257220P Ρ US 2000-257221P P 20001221 DE 2001-10111058 Α 20010308 DE 2001-10113366 Α 20010320 P US 2001-281653P 20010405 P US 2001-281874P 20010405 DE 2001-10138272 Α 20010810 US 2001-314599P P 20010824 US 2001-7182 B1 20011019 US 2001-86145 B1 20011019 US 2001-27662 B1 20011220 DE 2002-10206505 Α 20020216. A1 20020306 US 2002-92116 US 2002-93240 B1 20020307 US 2002-100659 A1 20020318 US 2002-369213P Ρ 20020401 US 2003-360064 A2 20030207 US 2003-413065 B2 20030414 US 2003-419358 A1 20030421 US 2003-613783 A2 20030703 US 2004-763894 A2 20040123 US 2004-775901 A2 20040210 US 2004-776757 A2 20040211 US 2004-824391 A2 20040414

OTHER SOURCE(S): MARPAT 138:44711

AB The present invention relates to novel pharmaceutical compns. based on anticholinergics and phosphodiesterase (PDE) IV inhibitors, processes for preparing them and their use in the treatment of respiratory tract diseases. For example, a suspension aerosol contained tiotropium bromide 0.029%, AWD 12-281 0.033%, ethanol 0.5%, iso-Pr myristate 0.1%, and TG 227 to 100%.

257892-33-4, AWD 12-281
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (inhalation compns. based on anticholinergics and phosphodiesterase IV
 inhibitors for treatment of respiratory tract diseases)

RN 257892-33-4 CAPLUS

IT

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

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CAPLUS COPYRIGHT 2006 ACS on STN
L25 ANSWER 29 OF 40
                         2002:695761 CAPLUS
ACCESSION NUMBER:
                         137:237718
DOCUMENT NUMBER:
                         Inhalant compositions containing anticholinergics and
TITLE:
                         PDE IV inhibitors
                         Meade, Christopher John Montague; Pairet, Michel;
INVENTOR(S):
                         Pieper, Michael Paul
                         Boehringer Ingelheim Pharma K.-G., Germany
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 34 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         German
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                DATE
                                           APPLICATION NO.
     PATENT NO.
                         KIND
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                               _____
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                                                                   _____
     WO 2002069945
                         A2
                                20020912
                                           WO 2002-EP1988
                                                                   20020226
     WO 2002069945
                         A3
                                20030130
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
             CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU; MC, NL, PT, SE, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                         DE 2001-10110772
     DE 10110772
                          Α1
                                20020912
                                                                   20010307
                                           CA 2002-2439763
     CA 2439763
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                                                                   20020226
                                          EP 2002-727329
     EP 1372649
                         A2
                                20040102
                                                                   20020226
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     JP 2004521134
                         T2
                                20040715
                                            JP 2002-569122
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     BR 2002007883
                         Α
                                20040727
                                            BR 2002-7883
                                                                   20020226
     NZ 528621
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                                            NZ 2002-528621
                                                                   20020226.
                                            CN 2002-805346
     CN 1649588
                         Α
                                20050803
                                                                   20020226
     ZA 2003006221
                         Α
                                20040722
                                            ZA 2003-6221
                                                                   20030812
PRIORITY APPLN. INFO.:
                                            DE 2001-10110772
                                                                A 20010307
                                            WO 2002-EP1988
                                                                W 20020226
OTHER SOURCE(S):
                         MARPAT 137:237718
     The invention relates to drug compns. based on anticholinergics and PDE IV
     inhibitors, to methods for their production, and to their use as inhalants for
     the treatment of respiratory tract diseases. Thus an inhalation powder
     was composed of capsules that contained (µg/capsule): tiotropium
     bromide 21.7; AWD-12-281 200; lactose 4778.3.
     257892-33-4, AWD-12-281
IT
     RL: PEP (Physical, engineering or chemical process); PYP (Physical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (inhalant compns. containing anticholinergics and PDE IV inhibitors)
RN
     257892-33-4 CAPLUS
     1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-
CN
     fluorophenyl) methyl]-5-hydroxy-α-οxo- (9CI) (CA INDEX NAME)
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Saloni Sharma 05/23/2006

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L25 ANSWER 30 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2002:575737 CAPLUS

DOCUMENT NUMBER:

137:135500

TITLE:

Methods of inducing ovulation by administering a

non-polypeptide cAMP level modulator

INVENTOR (S):

Palmer, Stephen; McKenna, Sean; Tepper, Mark; Eshkol,

Aliza; MacNamee, Michael C.

PATENT ASSIGNEE(S):

Applied Research Systems Holding N.V., USA

SOURCE:

U.S. Pat. Appl. Publ., 26 pp., Cont.-in-part of U.S.

Ser. No. 928,268.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

	PAT	CENT.	NO.			KINI)	DATE		AP	PL]	[CAT]	ON N	Ο.		DATE	E
•	US	2002	1031	06		A1	-	2002	0801	US	20	001-1	4812			2001	1214
	US	6953	774			B2		2005	1011								
	US	2002	0653	24		A1		2002	0530	US	20	001-9	92826	8		2001	10810
	CA	2469	939		•	AA		2003	0626	CA	. 20	001-2	24699	39		2001	1214
	AU	2002	2171	11		A1		2003	0630	AU	20	002-2	21711	1		2001	1214
	ΕP	1463	493			A1		2004	1006	EP	20	001-2	27498	7		2001	1214
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			IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY, A	L,	TR			•	•	
	BR	2001	0171	98 🍦		Α		2004	1026	BR	20	001-1	L7198			2001	1214
	CN	1582	146			Α		2005	0216	·CN	20	001-8	32395	1		2001	1214
	JP	2005	51692	24		T2		2005	0609	JP	20	003-5	55227	7		2001	1214
	US	2005	1485	01		A1		2005	0707	US	20	003-4	19863	9		2001	1214
	US	2006	0039	25		A 1		2006	0105	US	20	005-1	L6918	3		2005	0628
PRIO	IORITY APPLN. INFO.:									· US	20	000-2	22496	2 P	P	2000	0811
								-		US	20	001-9	92826	8	A2	2001	10810
										US	20	001-1	14812		A3	2001	L1214
										WC	20	001-E	EP147	30	W	2001	1214

AB The present invention relates to methods of inducing ovulation in a female host comprising the administration of a non-polypeptide cAMP level modulator to the female host. In another aspect, the invention provides for specific administration of the phosphodiesterase inhibitor prior to the luteal phase of the host's ovulatory cycle. Preferred non-polypeptide cAMP level modulator include phosphodiesterase inhibitors, particularly

inhibitors of phosphodiesterase 4 isoforms. Pharmaceutical compns. containing the cAMP modulators are also claimed.

IT 257892-33-4, AWD-12-281

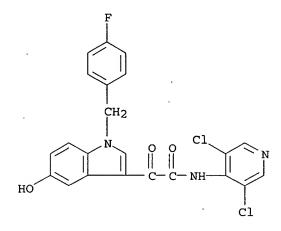
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(methods of inducing ovulation by administering a non-polypeptide cAMP level modulator)

RN . 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)



L25 ANSWER 31 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:495906 CAPLUS

DOCUMENT NUMBER: 138:117605

DOCUMENT NUMBER. 138.117003

TITLE: Effects of the phosphodiesterase 4 inhibitors SB

207499 and AWD 12-281 on the inflammatory reaction in

a model of allergic dermatitis

AUTHOR(S): Baumer, Wolfgang; Gorr, Gilbert; Hoppmann, Joachim;

Ehinger, Andreas M.; Ehinger, Britt; Kietzmann,

Manfred

CORPORATE SOURCE: Toxicology and Pharmacy, Department of Pharmacology,

School of Veterinary Medicine, Hanover, 30559, Germany

SOURCE: European Journal of Pharmacology (2002), 446(1-3),

195-200

CODEN: EJPHAZ; ISSN: 0014-2999

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

AB The inhibitors of the phosphodiesterase 4, SB 207499 (cilomilast, c-4-cyano-4-(3-cyclopentyloxy-4-methoxyphenyl)-r-L-cyclohexane carboxylic acid) and AWD 12-281 (N-(3,5-dichloropyrid-4-yl)-[1-(4-fluorobenzyl)-5-hydroxyindole-3-yl]glyoxylic acid amide) were tested in a model of allergic dermatitis in mice. To obtain an allergic dermatitis, BALB/c mice were sensitized to toluene-2,4-diisocyanate. The allergic reaction was challenged by topical administration of toluene-2,4-diisocyanate onto the mice ears. Before challenge, two groups of mice were treated topically (ear skin) with SB 207499 or AWD 12-281. There was a significant ear swelling in toluene-2,4-diisocyanate-challenged mice ears 4, 8, 16, 24 and 48 h after challenge. SB 207499 and AWD 12-281 inhibited this swelling significantly 8, 16, 24 and 48 h after the challenge. For

biochem. parameters and histol., ears were sampled from mice sacrificed 4, 8 and 16 h after the challenge. In homogenized tissue, SB 207499 and AWD 12-281 inhibited significantly the secretion of interleukin 1 β induced by toluene-2,4-diisocyanate 4 and 8 h after challenge. The cell influx (granulocytes) observed in the toluene-2,4-diisocyanate-challenged mice 8 and 16 h after challenge was nearly abolished by AWD 12-281 and SB 204799.

IT 257892-33-4, AWD 12-281

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(effects of phosphodiesterase 4 inhibitors SB 207499 and AWD 12-281 on inflammatory reaction in a model of allergic dermatitis)

RN 257892-33-4 CAPLUS

CN

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

REFERENCE COUNT: 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L25 ANSWER 32 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2001:850920 CAPLUS

DOCUMENT NUMBER:

135:366766

TITLE:

Method for enhancing cognitive function with

phosphodiesterase-4 inhibitors

INVENTOR(S):

Hagan, James

PATENT ASSIGNEE(S):

Smithkline Beecham P.L.C., UK

SOURCE:

PCT Int. Appl., 20 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT :	PATENT NO.					DATE		;	APPL	ICAT:	ION I	NO.		D	ATE	
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WO 2001	WO 2001087281					2001	1122	1	WO 2	001-	GB21	34.		2	0010	515
WO 2001	WO 2001087281					2002	0328			•						
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RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG EP 1292287 A2 20030319 EP 2001-929824 20010515 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR JP. 2003533473 T2 20031111 JP 2001-583749 20010515 US 2003187006 **A1** 20031002 US 2003-275853 20030314 PRIORITY APPLN. INFO.: GB 2000-11802 A 20000516 WO 2001-GB2134 W 20010515

A method for enhancing cognitive function by administering to a patient in need thereof an effective amount of a PDE4 inhibitor.

IT 257892-33-4, AWD-12-281

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(enhancing cognitive function with phosphodiesterase-4 inhibitors)

RN257892-33-4 CAPLUS .

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-CNfluorophenyl) methyl] -5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

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L25 ANSWER 33 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2001:380415 CAPLUS

DOCUMENT NUMBER:

134:361385

TITLE:

Combined phosphodiesterase 3 (PDE3) and

phosphodiesterase 4 (PDE4) inhibitor therapy for the

treatment of obesity

INVENTOR(S):

Snyder, Peter

PATENT ASSIGNEE(S):

Icos Corporation, USA PCT Int. Appl., 30 pp.

SOURCE: CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

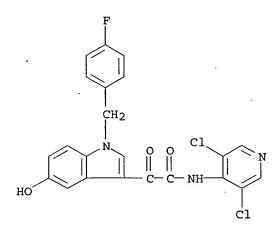
English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2000-US42137 WO 2001035979 A2 20010525 20001113 WO 2001035979 **A3** 20020103 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US. UZ. VN. YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM . RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG PRIORITY APPLN. INFO.: US 1999-165418P 19991113 Materials and methods are provided for the treatment of obesity that involve a combination of a PDE3 and PDE4 inhibitor in synergistically effective amts. Methods for producing PDE proteins are also described. IT 257892-33-4, AWD-12-281 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (phosphodiesterase 3 and phosphodiesterase 4 inhibitor combination therapy for treatment of obesity) 257892-33-4 CAPLUS RN CN1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl)methyl]-5-hydroxy-α-οάο- (9CI) (CA INDEX NAME)



L25 ANSWER 34 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2001:260010 CAPLUS

DOCUMENT NUMBER:

TITLE:

Requirement of additional adenylate cyclase activation for the inhibition of human eosinophil degranulation

by phosphodiesterase IV inhibitors

AUTHOR (S):

Ezeamuzie, C. I.

CORPORATE SOURCE:

Department of Pharmacology and Toxicology, Faculty of Medicine, P.O. Box 24923, Kuwait University, Safat,

13110, Kuwait

SOURCE:

European Journal of Pharmacology (2001), 417(1/2),

11-18

CODEN: EJPHAZ; ISSN: 0014-2999

PUBLISHER:

Elsevier Science B.V.

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Human eosinophils contain predominantly phosphodiesterase type IV, but selective inhibitors of this isoenzyme fail to inhibit certain eosinophil responses such as degranulation. In this study, the effect of activation of adenylate cyclase on the ability of several highly selective PDE IV inhibitors to inhibit complement C5a-induced O2- release and degranulation of human eosinophils in vitro was investigated. All four selective PDE IV inhibitors, N-(3,5-dichloropyrid-4-yl)-3-cyclopentyl-oxy-4methoxybenzamide (RP 73401), rolipram, N-(3,5-dichloropyrid-4-y1)-[1-(4fluorobenzyl)-5-hydroxy-indol-3-yl]glyoxylacidamide (AWD 12-281) and c-4-cyano-4-(3-cyclopentyloxy-4-methoxyphenyl-r-1-cyclohexane carboxylic acid) (SB 207499) potently inhibited C5a-induced O2- generation (IC50=0.03, 0.42, 0.55 and 0.86 μM , resp.), but generally failed to inhibit degranulation. The only exception was AWD 12-281, which inhibited degranulation (IC50=16.2 µM). In the presence of different AC activators (histamine, salbutamol, prostaglandin E2 and forskolin), the PDE IV inhibitors became potent inhibitors of degranulation. interaction between the PDE IV inhibitors and the AC activators resulted in a synergistic increase in intracellular levels of adenosine 3', 5'-monophosphate (cAMP). These results show that PDE IV inhibitors generally require an addnl. cAMP signal to be able to inhibit eosinophil degranulation, and that this signal can be generated via both membrane receptors and direct AC activation. This may be relevant to the in vivo effectiveness of PDE IV inhibitors in eosinophilic inflammation.

IT 257892-33-4, AWD 12-281

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(requirement of addnl. adenylate cyclase activation for inhibition of human eosinophil degranulation by phosphodiesterase IV inhibitors)

RN 257892-33-4 CAPLUS

CN

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo-(9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\$$

REFERENCE COUNT:

34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 35 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2001:259980 CAPLUS

DOCUMENT NUMBER:

135:57779

TITLE:

Identification of inhibitor binding sites of the

cAMP-specific phosphodiesterase 4

AUTHOR(S): Richter, W.; Unciuleac, L.; Hermsdorf, T.; Kronbach,

T.; Dettmer, D.

CORPORATE SOURCE: Medical Faculty, Institute of Biochemistry, University

of Leipzig, Leipzig, D-04103, Germany

SOURCE: Cellular Signalling (2001), 13(4), 287-297

CODEN: CESIEY; ISSN: 0898-6568

PUBLISHER: Elsevier Science Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

Using the technique of site-directed mutagenesis, point mutants of human PDE4A have been developed in order to identify amino acids involved in inhibitor binding. Relevant amino acids were selected according to a peptidic binding site model for PDE4 inhibitors, which suggests interaction with two tryptophan residues, one histidine and one tyrosine residue, as well as one Zn2+ ion. Mutations were directed at those tryptophan, histidine, and tyrosine residues, which are conserved among the PDE4 subtypes (PDE4A-D) and lie within the high-affinity 4-[3-(cyclopentoxyl)-4-methoxyphenyl]-2-pyrrolidone (rolipram) binding domain of human PDE4A (amino acids 276-681 according to the PDE4A sequence L20965). Truncations to this region do not alter enzyme activity or inhibitor sensitivity. The mutants were expressed in COS1 cells, and the recombinant cyclic nucleotide phosphodiesterase (PDE) forms have been characterized in terms of their catalytic activity and inhibitor sensitivities. Tyrosine residues 432 and 602, as well as histidine 588, were found to be involved in inhibitor binding, but no interaction was detected between tryptophan and PDE inhibitors tested. To test the possibility that other amino acids are of importance for hydrophobic interactions, selected phenylalanine residues were also mutated. We found phenylalanine 613 and 645 to influence inhibitor binding to PDE4. The significant differences in the inhibitor sensitivities of the mutants show that the various inhibitors have different enzyme binding sites. Based on the assumption that the known side effects of PDE4 inhibitors (like emesis and nausea) are caused directly by selective inhibition of different conformation states of PDE4, our results may be a hint to differ between PDE4 inhibitors, which have emetic side effects (like rolipram), and those that do not have side effects (like N-(3,5-dichlorpyrid-4-yl)-[1-(4fluorbenzyl)-5-hydroxy-indol-3-yl]-glyoxylateamide [AWD12-281]) by the differences of their binding sites and in that context contribute to the development of novel drugs. Furthermore, the identification of amino acid interactions proposed by the peptidic binding site model, which was used for the mutant selection, verifies the PrGen modeling as a useful method for the prediction of inhibitor binding sites in cases where detailed knowledge of the protein structure is not available.

IT **257892-33-4**, AWD12-281

RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (identification of inhibitor binding sites of cAMP-specific phosphodiesterase 4)

RN 257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

Saloni Sharma 05/23/2006

$$\begin{array}{c|c} F \\ \hline \\ CH_2 \\ \hline \\ N \\ \hline \\ C \\ C \\ C \\ \end{array}$$

REFERENCE COUNT: 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 36 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:30560 CAPLUS

DOCUMENT NUMBER: 134:221365

TITLE: The effect of selective and non-selective

phosphodiesterase inhibitors on allergen- and leukotriene C4-induced contractions in passively

sensitized human airways

AUTHOR(S): Schmidt, Dunja T.; Watson, Nikki; Dent, Gordon;

Ruhlmann, Elke; Branscheid, Detlev; Magnussen, Helgo;

Rabe, Klaus F.

CORPORATE SOURCE: Department of Pulmonology, Leiden University Medical

Centre, Leiden, NL-2333 ZA, Neth.

SOURCE: British Journal of Pharmacology (2000), 131(8),

1607-1618

CODEN: BJPCBM; ISSN: 0007-1188

PUBLISHER: Nature Publishing Group

DOCUMENT TYPE: Journal LANGUAGE: English

Non-selective inhibitors of cyclic nucleotide phosphodiesterase (PDE) block allergen-induced contraction of passively sensitized human airways in vitro by a dual mechanism involving a direct relaxant effect on smooth muscle and inhibition of histamine and cysteinyl leukotriene (LT) release from airways. We investigated the effects of non-selective PDE inhibitors and selective inhibitors of PDE3 and PDE4 in order to determine the involvement of PDE isoenzymes in the suppression of allergic bronchoconstriction. Macroscopically normal airways from 76 patients were sensitized with IgE-rich sera (>250 u ml-1) containing specific antibodies against allergen (Dermatophagoides farinae). Contractile responses of bronchial rings were assessed using standard organ bath techniques. Passive sensitization caused increased contractile responses to allergen, histamine and LTC4. Non-selective PDE inhibitors (theophylline, 3-isobutyl-1-methylxanthine [IBMX]), a PDE3-selective inhibitor (motapizone), PDE4-selective inhibitors (RP73401, rolipram, AWD 12-281) and a mixed PDE3/4 inhibitor (zardaverine) all significantly relaxed inherent bronchial tone at resting tension and to a similar degree. Theophylline, IBMX, zardaverine and the combination of motapizone and RP73401 inhibited the contractile responses to allergen and LTC4. Pre-treatment with motapizone, RP73401, rolipram or the methylxanthine adenosine receptor antagonist, 8-phenyltheophylline, did not significantly decrease responses to either allergen or LTC4. We

conclude that combined inhibition of PDE3 and PDE4, but not selective inhibition of either isoenzyme or antagonism of adenosine receptors, is effective in suppressing allergen-induced contractions of passively sensitized human airways. The relationship between allergen- and LTC4-induced responses suggests that PDE inhibitors with PDE3 and PDE4 selectivity are likely to act in part through inhibition of mediator release and not simply through direct relaxant actions on airway smooth muscle.

257892-33-4, AWD 12-281

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)

(phosphodiesterase inhibitors in allergen- and leukotriene C4-induced contractions in sensitized human airways)

RN 257892-33-4 CAPLUS

1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4-fluorophenyl)methyl]-5-hydroxy-α-oxo-(9CI) (CA INDEX NAME)

REFERENCE COUNT: 52 THERE ARE 52 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 37 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1999:647583 CAPLUS

DOCUMENT NUMBER:

132:145941

TITLE:

CN

Therapeutic potential of phosphodiesterase 4

inhibitors in allergic diseases

AUTHOR(S):

Crocker, I. Caroline; Townley, Robert G.

CORPORATE SOURCE:

Creighton University Allergic Disease Center, Omaha,

NE, USA

SOURCE:

Drugs of Today (1999), 35(7), 519-535

CODEN: MDACAP; ISSN: 0025-7656

PUBLISHER:

Prous Science

DOCUMENT TYPE:

Journal; General Review

LANGUAGE: English

AB A review with 137 refs. CAMP is thought to be associated with inflammatory cell activity: high levels tend to decrease proliferation and cytokine secretion, whereas low concns. have the opposite effect (1). Since many phosphodiesterases (PDEs) degrade cAMP, inhibitors of this enzyme decrease inflammatory cell activity. Theophylline, which has nonselective PDE inhibitor activity in addition to its other mechanisms of action, has been used in the treatment of asthma for many years. Unfortunately, because of the important role of PDEs in the cell, nonspecific inhibition of these

enzymes causes many undesirable side effects. The discovery of PDE isoenzyme families (PDE1-PDE10), their subtypes (HPDE4 and LPDE4) and their differential distribution among the cell types, as well as their specific functions in controlling cell processes, has led to the development of new, specific PDE4 inhibitors. This review details the rationale for the use of PDE4 inhibitors in the treatment of allergic disease. In addition, the effects of PDE4 inhibitors in vitro, in preclin. animal models and in the clinic are covered. Finally, up-to-date information on the most recently developed inhibitors, such as SB-207499, CDP-840, AWD-12-281 and D-4418, is provided.

ΙT 257892-33-4, AWD 12-281

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(therapeutic potential of phosphodiesterase 4 inhibitors in allergic diseases)

RN257892-33-4 CAPLUS

CN 1H-Indole-3-acetamide, N-(3,5-dichloro-4-pyridinyl)-1-[(4fluorophenyl) methyl] -5-hydroxy- α -oxo- (9CI) (CA INDEX NAME)

REFERENCE COUNT:

137 THERE ARE 137 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

L25 ANSWER 38 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN .

ACCESSION NUMBER: 1998:175908 CAPLUS

DOCUMENT NUMBER: 128:217285

TITLE: Preparation of new, N-substituted indole-3-

glyoxylamides as antiasthmatics, antiallergic agents

and immunosuppressants/immunomodulators

INVENTOR(S): Lebaut, Guillaume; Menciu, Cecilia; Kutscher,

Bernhard; Emig, Peter; Szelenyi, Stefan; Brune, Kay

PATENT ASSIGNEE(S): Asta Medica Aktiengesellschaft, Germany

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND APPLICATION NO. DATE DATE

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WO	9809946	_		A1		1998	0312		wo	1997-	EP4	174	-	-	9970	816	
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AU	9740158	}		A1		1998	0326		AU	1997	-401	58		-	19970	816	
AU	726521	•		B2		2000	1109							•		010	
EP	726521 931063			A1		1999	0728		ΕP	1997	-937	586		•	19970	816	
	R: AT	BE.	CH.	DE.	DK.	ES.	FR.	GB.	GR	IT.	LI	. LU.	NL.	SE	MC.	PT.	
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CN	1227542	·		Α		1999	0901		CN	1997	-197	128		:	19970	816	
BR	9712808	3		Α		1999	1123		BR	1997	-128	80		:	19970	816	
JP	2000509	098		T2		2000	0425		JP	1998	-512	167		:	19970	816	
JР	3296437	7		B2		2002	0702										
NZ	334476			A	•	2000	0526		NZ	1997	-334	476		-	19970	816	
IL	127798			A1		2003	0731		IL	1997	-127	798			19970	816	
. CN	1496980)		Α		2004	0519		CN	2002	-200	21320	61	:	19970	816	
RU	2237661	t		C2		2004	1010		RU	1999	-106	782		:	19970	816	
ZA	9707475	5 \		Α		1998	0219		ZA	1997	-747	5	•	-	19970	820	
CA	2215013	3 .		AA		1998	0306		CA	1997	-221	5013		-	19970	904	
· CA	2215013	3		C													
US	6008231			Α								326					
TW	550256			В		2003	0901		TW	1997	-861	12985		-	19970	930	
NO	9901071	L		Α		1999	0304		NO	1999	-107	1		-	19990	304	
NO	2215013 2215013 6008231 550256 9901071 314725 6344467			B1			0512										
US	6344467	7		B1		2002	0205		US	1999	-409	263			19990	930	
US	2002161	L025		A 1		2002	1031		US	2002	-588	36		2	20020	130	
NO	2003000	0481		Α		1999	0304		NO	2003	-481			2	20030	130	
US	2002161 2003000 2003203 6919344	7892		A1		2003	1106		US	2003	-402	36 931		2	20030	401	
US	6919344	<u> </u>		B2		2005	0719										
PRIORITY	Y APPLN	INFO).:						DE	1996	-196	36150 474 326 263		A :	19960	906	
									WO	1997	-EP4	474		W :	19970	816	
									US	1997	-925	326		A3 :	19970	908	
									US	1999	-409	263		A3 :	19990	930	
									US	2002	-588	36		B1 2	20020	130	
								~ -									

OTHER SOURCE(S):

MARPAT 128:217285

GΙ

$$\begin{array}{c|cccc}
R^4 & & Z & R & R^1 \\
\hline
R^3 & & R^2 & & I
\end{array}$$

The title compds. [I; R = H, (un)substituted C1-6 alkyl; R1 = (un)substituted Ph, pyridyl, pyrimidinyl, etc.; RR1 = atoms to close (N-substituted) piperazine ring; R2 = H, (un)substituted C1-6 alkyl, (un)substituted benzoyl; R3, R4 = H, OH, C1-6 alkyl, C3-7 cycloalkyl, halo, NO2, amino, benzyloxy, etc.; Z = O, S] and their acid salts were prepared, e.g., by N-alkylation of indoles with R2-bearing reactants followed by acylation with a dicarbonyl halide and amidation of the remaining acid halide function. For example, a title compound I (R = R3 = R4 = H, R1 = 4-pyridyl, R2 = 4-FC6H4CH2, Z = O) (preparation by benzylation of

Saloni Sharma

indole with 4-FC6H4CH2Cl, acylation of the intermediate with (COCl)2 and amidation of the acyl chloride with 4-aminopyridine given) at 10 mg/kg i.p. in guinea pigs gave 55.4% inhibition of allergen-induced late-phase eosinophilia, vs. 47.0 for cyclosporin A.

IT 204206-02-0P

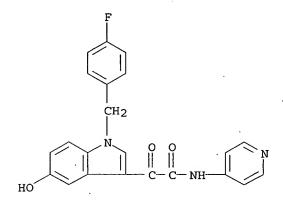
CN

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of N-substituted indoleglyoxylamides as antiasthmatics, antiallergic agents and immunosuppressants/immunomodulators)

RN 204206-02-0 CAPLUS

1H-Indole-3-acetamide, 1-[(4-fluorophenyl)methyl]-5-hydroxy- α -oxo-N-4-pyridinyl- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L25 ANSWER 39 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

5

ACCESSION NUMBER:

1992:571228 CAPLUS

DOCUMENT NUMBER:

117:171228

TITLE:

Preparation of N-(azabicycloalkyl)indole-3-glyoxylamides and analogs as 5-HT antagonists

INVENTOR (S):

Clark, Robin D.; Eglen, Richard M.; Muchowski, Joseph

M.; Smith, William L.; Weinhardt, Klaus K.

PATENT ASSIGNEE(S):

SOURCE:

Syntex (U.S.A.), Inc., USA Eur. Pat. Appl., 38 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent English

1

LANGUAGE:
FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.		KIND	DATE	APPLICATION NO.	DATE
	EP 490263		A1	19920617	EP 1991-120856	19911204
	R: AT	, BE, CH,	DE, DE	C, ES, FR,	GB, GR, IT, LI, LU, I	NL, SE
_	US 5192770		A	19930309	US 1990-624028	19901207
	FI 9105736		A	19920608	FI 1991-5736	19911205
	CA 2057181		AA	19920608	CA 1991-2057181	19911206
	NO 9104825		Α	19920609	NO 1991-4825	19911206
	AU 9188856	•	A1	19920611	AU 1991-88856	19911206
	AU 644249		B2	19931202		
	HU 60270		A 2	19920828	HU 1991-3836	19911206

JP 04290884 A2 19921015 JP 1991-322979 19911206
ZA 9109660 A 19930607 ZA 1991-9660 19911206
PRIORITY APPLN. INFO.: US 1990-624028 A 19901207
OTHER SOURCE(S): MARPAT 117:171228

AB R1COCONR2R3 [I; R1 = (substituted) Ph, -indolo, -2-oxobenzimidazolo, -3-benzofuranyl, -3-indolyl, etc.; R2 = azabicycloalkyl groups Q1, Q2, etc.; R3 = H, alkyl; R6 = alkyl; p = 0, 1; q = 1-3] were prepared Thus, 1-methyl-α-oxo-3-indoleacetyl chloride was condensed with (S)-3-amino-1-azabicyclo[2.2.2]octane to give title compound (S)-II which reversed atropine-induced cognitive deficit in mice at .apprx.1 mg/kg orally.

IT 143137-41-1P 143137-42-2P 143137-43-3P 143339-42-8P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of, as 5-HT antagonist)

RN 143137-41-1 CAPLUS

GI

CN lH-Indole-3-acetamide, N-1-azabicyclo[2.2.2]oct-3-yl-5-hydroxy-1-methyl- α -oxo- (9CI) (CA INDEX NAME)

RN 143137-42-2 CAPLUS

CN 1H-Indole-3-acetamide, N-1-azabicyclo[2.2.2]oct-3-yl-5-hydroxy-1-methyl- α -oxo-, (S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 143137-43-3 CAPLUS

CN 1H-Indole-3-acetamide, N-1-azabicyclo[2.2.2]oct-3-yl-5-hydroxy-1-methyl- α -oxo-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN143339-42-8 CAPLUS

1H-Indole-3-acetamide, N-1-azabicyclo[2.2.2]oct-3-yl-5-hydroxy-1-methyl- α -oxo-, monohydrochloride, (S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

HC1

L25 ANSWER 40 OF 40 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1969:413112 CAPLUS

DOCUMENT NUMBER: 71:13112

Recent progress in rifamycin derivatives chemistry TITLE:

AUTHOR(S): Maggi, Nicola; Sensi, Piero

CORPORATE SOURCE: Res. Lab., Lepetit S.p.A., Milan, Italy

SOURCE: Int. Congr. Chemother., Proc., 5th (1967), Volume 1, Issue 1, 15-21. Editor(s): Spitzy, K. H. Verlag

Wien. Med. Akad.: Vienna, Austria.

CODEN: 20JJA4

DOCUMENT TYPE: Conference LANGUAGE: English

GI For diagram(s), see printed CA Issue.

AB Cf. CA 65: 12597c. An investigation of derivs. of 3-formylrifamycin SV (I) bearing amino groups on the side chain is reported. O-substituted oximes, N-substituted imines and hydrazones of I are cited. None is as active on Staphylococcus aureus as Rifampicin. A new series of rifamycins with a pyrrole nucleus condensed in positions 3 and 4 is also reported. These products are obtained by reaction of rifamycin S with allyl amines. The structure of the resulting 3,4-pyrrolorifamycins (II) was confirmed by N.M.R. and uv. Bacterial activity is dependent upon size of substituents R, R1, and R2. In the presence of Et3N the same reaction yields products with a hydroquinone structure (III). The N.M.R. spectrum shows the presence of a methinic hydrogen.

IT 22912-69-2P

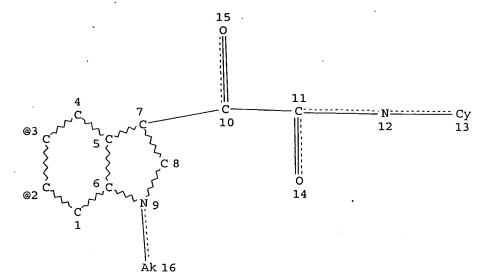
RN 22912-69-2 CAPLUS

CN 9,4-(Epoxypentadeca[1,11,13]trienimino)-1H-benzofuro[5,4-g]indole-3-glyoxylanilide, 9,10-dihydro-5,6,16,18,20-pentahydroxy-14-methoxy-1,2,7,9,15,17,19,21,25-nonamethyl-10,26-dioxo-, 16-acetate (8CI) (CAINDEX NAME)

d que 124 74) SEA FILE=CAPLUS ABB=ON L1 PLU=ON ("RUNDFELDT C"/AU OR "RUNDFELDT CHRIS"/AU) L251) SEA FILE=CAPLUS ABB=ON PLU=ON ("KUSS H"/AU OR "KUSS H H"/AU OR "KUSS H J"/AU OR "KUSS H M"/AU OR "KUSS HILDEGARD"/AU) 12) SEA FILE=CAPLUS ABB=ON L3 PLU=ON ("HOFGEN N"/AU OR "HOFGEN NORBERT"/AU) L4 9 SEA FILE=CAPLUS ABB=ON PLU=ON (L1 AND (L2 OR L3)) OR (L2 AND L3) L5 74) SEA FILE=CAPLUS ABB=ON PLU=ON ("RUNDFELDT C"/AU OR "RUNDFELDT CHRIS"/AU) L6 51) SEA FILE=CAPLUS ABB=ON PLU=ON ("KUSS H"/AU OR "KUSS H H"/AU OR "KUSS H J"/AU OR "KUSS H M"/AU OR "KUSS HILDEGARD"/AU) 12) SEA FILE=CAPLUS ABB=ON PLU=ON ("HOFGEN N"/AU OR "HOFGEN L7

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NORBERT"/AU)
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L8 126 SEA FILE=CAPLUS ABB=ON PLU=ON (L5 OR L6 OR L7) L16 STR



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NODE ATTRIBUTES:
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IS M1

NSPEC IS R ΑT 1 NSPEC IS R AΤ 2 IS R NSPEC AΤ 3 NSPEC IS R AΤ NSPEC IS R AT 5 IS R NSPEC TA6 NSPEC IS R TA7 NSPEC IS R AΤ 8 NSPEC IS R AT. 9 **NSPEC** IS C ΤA 10 NSPEC IS C ΑT 11 NSPEC IS C AΤ 12 NSPEC IS C TA13 NSPEC IS C AT14 NSPEC IS C AΤ 15 NSPEC IS C AΤ 16 IS C NSPEC AΤ 17 DEFAULT MLEVEL IS ATOM

MLEVEL IS CLASS AT 10 11 12 14 15 16 17

AΤ

17

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 17

STEREO ATTRIBUTES: NONE

L19 35 SEA FILE=REGISTRY SSS FUL L16
L20 51 SEA FILE=CAPLUS ABB=ON PLU=ON L19

L22 11 SEA FILE=CAPLUS ABB=ON PLU=ON L20 AND L8 L24 14 SEA FILE=CAPLUS ABB=ON PLU=ON (L4 OR L22)

=> d ibib abs 124 tot

AUTHOR (S):

L24 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:226501 CAPLUS

DOCUMENT NUMBER: 144:267237

TITLE: The phosphodiesterase 4 inhibitor AWD 12-281 is active

in a new guinea-pig model of allergic skin

inflammation predictive of human skin penetration and

suppresses both Th1 and Th2 cytokines in mice Hoppmann, Joachim; Baeumer, Wolfgang; Galetzka, Christin; Hoefgen, Norbert; Kietzmann, Manfred;

Rundfeldt, Chris

CORPORATE SOURCE: Department of Pharmacology, elbion AG, Radebeul,

D-01445, Germany

SOURCE: Journal of Pharmacy and Pharmacology (2005), 57(12),

1609-1617

CODEN: JPPMAB; ISSN: 0022-3573

PUBLISHER: Pharmaceutical Press

DOCUMENT TYPE: Journal LANGUAGE: English

The selective phosphodiesterase 4 (PDE4) inhibitor AWD 12-281 is structurally optimized for topical administration. It has potent effects in models of lung inflammation if administered as a dry powder inhalation. It has also demonstrated its anti-inflammatory property in a mouse model of cutaneous inflammation after topical administration. The aim of this study was to evaluate whether AWD 12-281 may be capable of penetrating human skin. Therefore a new guinea-pig model of allergic skin inflammation had to be developed. In ovalbumin-sensitized quinea-pigs, intracutaneous administration of ovalbumin results in a rapid development of allergic skin wheals. Topically administered AWD 12-281 was capable of reducing the development of wheals, indicating that this compound can penetrate the stratum corneum of quinea-piq skin as a predictor of human skin penetration. A secondary aim was the evaluation of a T cell subtype preference of AWD 12-281 since PDE4 inhibitors are said to preferentially inhibit Th2-type cytokines. Therefore, the effects of AWD 12-281 on a broad spectrum of Th1- and Th2-type cytokines were studied in tissue homogenates after allergen challenge in sensitized mice and in supernatants of anti CD3/anti-CD28-stimulated peripheral blood mononuclear cells (PBMCs). In both models, AWD 12-281 suppressed both T cell subtype cytokines indicating a broad spectrum activity of AWD 12-281. A further issue was to determine the duration of action and the concentration-response relation

of the topical activity of AWD 12-281 using a model of acute local inflammation - the arachidonic-acid-induced mouse ear edema. The compound exhibited a dose-dependent effect with a minimally effective concentration of 0.3%; after repeated administration the minimally effective concentration was 0.03%. A single administration of a 3% solution resulted in significant suppression of inflammation even 48 h after treatment. In conclusion, our results indicate that AWD 12-281 is a very promising drug candidate not only for the treatment of lung inflammation using inhalative administration but also for the treatment of atopic dermatitis.

REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:927204 CAPLUS

DOCUMENT NUMBER: 141:395538

TITLE: Preparation of 7-azaindolylglyoxylamides as

phosphodiesterase IV inhibitors.

INVENTOR(S):

Hoefgen, Norbert; Kuss, Hildegard; Olbrich, Matthias; Egerland, Ute; Rundfeldt, Chris;

Steinike, Karin; Schindler, Rudolf

PATENT ASSIGNEE(S): SOURCE:

Elbion A.-G., Germany PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent German

LANGUAGE:

Germa

Ι

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 2004094416		WO 2004-EP4339	20040423
W: AE, AG, AL	, AM, AT, AU, AZ,	BA, BB, BG, BR, BW, B	Y, BZ, CA, CH,
		DM, DZ, EC, EE, EG, E	
GE, GH, GM	, HR, HU, ID, IL,	IN, IS, JP, KE, KG, K	P, KR, KZ, LC,
		MD, MG, MK, MN, MW, M	
		RO; RU, SC, SD, SE, S	
		UG, US, UZ, VC, VN, Y	
		SD, SL, SZ, TZ, UG, Z	
		AT, BE, BG, CH, CY, C	
		IT, LU, MC, NL, PL, P	
	, BJ, CF, CG, CI,	CM, GA, GN, GQ, GW, M	L, MR, NE, SN,
TD, TG			
	=	DE 2003-10318610	
		US 2004-826136	-
AU 2004232483	A1 20041104	AU 2004-232483	20040423
CA 2523063	AA 20041104	CA 2004-2523063	20040423
EP 1613627	A1 20060111	EP 2004-729102	20040423
R: AT, BE, CH		GB, GR, IT, LI, LU, N	
		CY, AL, TR, BG, CZ, E	
PRIORITY APPLN. INFO.:	•	DE 2003-10318610	
		WO 2004-EP4339	
OTHER SOURCE(S):	MARPAT 141:3955		

$$\begin{array}{c|c}
0 & R^2 \\
N & B \\
R^4
\end{array}$$

GI

Title compds. [I; A = N, N-oxide group; B = C, N, N-oxide group; R1 = (substituted) alkyl, alkenyl; R2 = H, alkyl; R3, R4 = H, alkyl, OH, SH, NH2, NO2, cyano, SO3H, CO2H, alkoxycarbonyl, halo, alkoxy, alkylthio, (substituted) Ph, pyridyl, etc.], were prepared Thus, N-(3,5-dichloropyridin-4-yl) [1-(4-fluorobenzyl)-7-azaindol-3-yl]glyoxylic acid amide in CH2Cl2 was treated dropwise with m-chloroperbenzoic acid in HOAc followed by stirring for 7 days to give 9.4% N-(3,5-dichloropyridin-4-yl) [1-(4-fluorobenzyl)-7-oxo-7-azaindol-3-yl]glyoxylic acid amide. I inhibited phosphodiesterase 4 with IC50's in the range of 10-10 M to 10-5 M.

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CAPLUS COPYRIGHT 2006 ACS on STN L24 ANSWER 3 OF 14

ACCESSION NUMBER: 2004:927194 CAPLUS

DOCUMENT NUMBER: 141:395426

TITLE:

Preparation of N-oxopyridinyl

hydroxyindolylglyoxylamides as phosphodiesterase IV

inhibitors.

INVENTOR (S): Hoefgen, Norbert; Kuss, Hildegard; Steinike,

Karin; Egerland, Ute; Rundfeldt, Chris;

Pfeifer, Thomas

PATENT ASSIGNEE(S):

Elbion A.-G., Germany PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent German

LANGUAGE:

SOURCE:

Ι

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION: DATERNITE NO.

PATENT NO.	KIND DATE	APPLICATION NO.	
WO 2004094406	A1 20041104	WO 2004-EP4340	
W: AE, AG, AI	, AM, AT, AU, AZ,	BA, BB, BG, BR, BW,	BY, BZ, CA, CH,
CN, CO, CF	t, CU, CZ, DE, DK,	DM, DZ, EC, EE, EG,	ES, FI, GB, GD,
GE, GH, GN	I, HR, HU, ID, IL,	IN, IS, JP, KE, KG,	KP, KR, KZ, LC,
LK, LR, LS	G, LT, LU, LV, MA,	MD, MG, MK, MN, MW,	MX, MZ, NA, NI,
NO, NZ, ON	I, PG, PH, PL, PT,	RO, RU, SC, SD, SE,	SG, SK, SL, SY,
TJ, TM, TN	I, TR, TT, TZ, UA,	UG, US, UZ, VC, VN,	YU, ZA, ZM, ZW
RW: BW, GH, GN	I, KE, LS, MW, MZ,	SD, SL, SZ, TZ, UG,	ZM, ZW, AM, AZ,
BY, KG, K2	K, MD, RU, TJ, TM,	AT, BE, BG, CH, CY,	CZ, DE, DK, EE,
. ES, FI, FF	, GB, GR, HU, IE,	IT, LU, MC, NL, PL,	PT, RO, SE, SI,
SK, TR, BI	r, BJ, CF, CG, CI,	CM, GA, GN, GQ, GW,	ML, MR, NE, SN,
TD, TG			
DE 10318609	A1 20041111	DE 2003-10318609	20030424
US 2004266760	A1 20041230	US 2004-824342	20040414
AU 2004232484	A1 20041104	AU 2004-232484	20040423
CA 2523062	AA 20041104	CA 2004-2523062	20040423
EP 1615911	A1 20060118	EP 2004-729060	20040423
R: AT, BE, CH	I, DE, DK, ES, FR,	GB, GR, IT, LI, LU,	NL, SE, MC, PT,
IE, SI, LI	C, LV, FI, RO, MK,	CY, AL, TR, BG, CZ,	EE, HU, PL, SK, HR
BR 2004009683	A 20060418	BR 2004-9683	20040423
PRIORITY APPLN. INFO.:		DE 2003-10318609	A 20030424
		WO 2004-EP4340	A 20040423
OTHER SOURCE(S):	MARPAT 141:3954	26	

AB Title compds. [I; R1 = (substituted) alkyl, alkenyl; R2 = H, alkyl; R3 = OH; R4, R5 = H, alkyl, OH, SH, NH2, NO2, cyano, SO3H, CO2H, alkoxycarbonyl, halo, alkoxy, alkylthio, (substituted) Ph, pyridyl, etc.], were prepared Thus, N-(3,5-dichloropyridin-4-yl) [5-benzyloxy-1-(4fluorobenzyl)indol-3-yl]glyoxylamide in CH2Cl2 was treated dropwise with m-chloroperbenzoic acid in HOAc followed by stirring for 7 days to give 16.1% pyridine N-oxide derivative, which was refluxed with BBr3 in CH2Cl2 to give 72.8% N-(3,5-dichloro-1-oxopyridin-4-yl) [1-(4-fluorobenzyl)-5hydroxyindol-3-yl]glyoxylamide. I inhibited phosphodiesterase 4 with IC50's in the range of 10-5 M to 10-10 M. REFERENCE COUNT: THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS

L24 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN .

ACCESSION NUMBER:

2004:927193 CAPLUS

DOCUMENT NUMBER:

141:395425

TITLE:

Preparation of hydroxyindolylglyoxylic acid

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT ·

oxopyridinylamides as phosphodiesterase IV inhibitors. INVENTOR(S): Hoefgen, Norbert; Kuss, Hildegard; Steinike,

Karin; Egerland, Ute; Rundfeldt, Chris

PATENT ASSIGNEE(S):

Elbion A.-G., Germany

SOURCE:

PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATI	ENT N	0.			KINI		DATE		;	APPL	ICAT	ION I	NO.		D	ATE		
WO 2	20040	9440					2004	1104	1	WO 2	004-	EP43	38		2	00404	423	
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	4	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
							ID,											
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,	
		NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	·SG,	SK,	SL,	SY,	
							TZ,											
	RW: BW, GH, GN BY, KG, K																	
	BY, KG, K																	
	ES, FI, F																	
				BF,	ВJ,	CF,	CG,	CI,	CM,	·GA,	GN,	GQ,	GW,	ML,	MR,	NΕ,	SN,	
		TD,																
	10318				A1		2004											
	20042						2004											
	20042						2004											
	25230				AA		2004											
EP :	16159						2006											
	R: AT, BE, CH																	
	IE, SI, LT,				LV,	FI,	RO,	MK,										HR
PRIORITY	ORITY APPLN. INFO.:											1031						
	ED GOVERGE (G)									WO 2	004-	EP43	38	1	W 20	00404	423	
OTHER SOU GI	ER SOURCE(S):				MAR	PAT	141:	3954:	25									

Saloni Sharma

Title compds. [I; R1 = (substituted) alkyl, alkenyl; R2 = H, alkyl; R3-R5 AB = H, OH; \geq 1 or R3-R5 = OH; R6, R7 = H, alkyl, OH, SH, NH2, NO2, cyano, SO3H, CO2H, alkylcarbonyloxy, halo, alkylthio, (substituted) Ph, pyridyl, etc.], were prepared Thus, N-(3,5-dichloropyridin-4-yl) [7-benzyloxy-1-(4-fluorobenzyl)indol-3-yl]glyoxylic acid amide was stirred 7 days with m-chloroperbenzoic acid in HOAc to give 16.9% pyridine N-oxide derivative, which was refluxed with BBr3 in CH2Cl2 to give 66.2% N-(3,5-dichloro-1-oxopyridin-4-yl) [1-(4-fluorobenzyl)-7-hydroxyindol-3yl]glyoxylic acid amide. I inhibited phosphodiesterase 4 with IC50's in the range of 10-10 M to 10-5 M.

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS 6 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2004:450478 CAPLUS

DOCUMENT NUMBER:

141:23423

Ι

TITLE:

Preparation of 4- and/or 7-hydroxyindoles as

phosphodiesterase 4 inhibitors

INVENTOR(S):

Hoefgen, Norbert; Kuss, Hildegard; Egerland, Ute; Rundfeldt, Chris; Hartenhauer, Helge;

Gasparic, Antje

PATENT ASSIGNEE(S):

SOURCE:

Elbion Ag, Germany Ger. Offen., 17 pp.

CODEN: GWXXBX Patent .

DOCUMENT TYPE:

German LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE				
DE 10253426	A1 200406	DE 2002-10253426	20021115				
DE 10253426	B4 200509:	22					
US 2004147759	A1 200407	29 US 2003-714568	20031113				
CA 2505988	AA 200406	03 CA 2003-2505988	20031114				
WO 2004045607	A1 200406	03 WO 2003-EP12742	20031114				
W: AE, AG,	AL, AM, AT, AU, A	Z, BA, BB, BG, BR, BW,	BY, BZ, CA, CH,				
· · · · · · · · · · · · · · · · · · ·	•	K, DM, DZ, EC, EE, EG,					
GE, GH,	GM, HR, HU, ID, II	L, IN, IS, JP, KE, KG,	KP, KR, KZ, LC,				
LK, LR,	LS, LT, LU, LV, M	A, MD, MG, MK, MN, MW,	MX, MZ, NI, NO,				
NZ, OM,	PG, PH, PL, PT, R	O, RU, SC, SD, SE, SG,	SK, SL, SY, TJ,				
TM, TN,	TR, TT, TZ, UA, U	G, US, UZ, VC, VN, YU,	ZA, ZM, ZW				
RW: BW, GH,	GM, KE, LS, MW, M	Z, SD, SL, SZ, TZ, UG,	ZM, ZW, AM, AZ,				
BY, KG,	KZ, MD, RU, TJ, T	M, AT, BE, BG, CH, CY,	CZ, DE, DK, EE,				
ES, FI,	FR, GB, GR, HU, I	E, IT, LU, MC, NL, PT,	RO, SE, SI, SK,				
TR, BF,	BJ, CF, CG, CI, C	M, GA, GN, GQ, GW, ML,	MR, NE, SN, TD, TG				
AU 2003283400	A1 200406	15 AU 2003-283400	20031114				

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EP 1562584
                                20050817
                          A1
                                             EP 2003-775355
                                                                     20031114
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     BR 2003016234 .
                          Α
                                20051011
                                             BR 2003-16234
                                                                     20031114
     JP 2006508141
                          T2
                                 20060309
                                             JP 2004-552596
                                                                     20031114
     NO 2005002864
                                20050613
                                             NO 2005-2864
                                                                     20050613
PRIORITY APPLN. INFO.:
                                             DE 2002-10253426
                                                                  A 20021115
                                             WO 2003-EP12742
                                                                    20031114
OTHER SOURCE(S):
                         MARPAT 141:23423
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$$\begin{array}{c|c}
R^5 & NR^1R^2 \\
\downarrow & \downarrow & \downarrow \\
NR^1 & (C=0)_n
\end{array}$$

GT

AB Title compds. [I; n = 1, 2; R1 = (substituted) (branched) alkyl, (substituted) (branched) unsatd. alkenyl; R2, R3 = H, (substituted) alkyl, pyridyl, etc.; R4, R5 = H, OH], were prepared Thus, a suspension of NaH in THF was dropwise treated with 4-amino-3,5-dichloropyridine in THF followed by stirring for 1 h at 20°. The reaction mixture was dropwise treated with 7-benzyloxy-1-(4-chlorobenzyl)-indol-3-ylglyoxyloyl chloride (preparation given) at 0° followed by reflux for 4 h to give 47.5% N-(3,5-dichloropyridin-4-yl)-[1-(4-chlorobenzyl)-7-hydroxyindol-3-yl]glyoxylamide. The latter inhibited phosphodiesterase 4 (PDE 4) with IC50 = 0.002 μmol/L.

L24 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:216863 CAPLUS

I

DOCUMENT NUMBER: 140:247052

TITLE: Treatment nonallergic rhinitis by selective

phosphodiesterase 4 inhibitors

INVENTOR(S): Rundfeldt, Chris; Kuss, Hildegard;

Hofgen, Norbert

PATENT ASSIGNEE(S): Elbion A.-G., Germany SOURCE: Ger. Offen., 12 pp.

CODEN: GWXXBX

CODEN: GWAXBA

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLIC	APPLICATION NO.							
DE 102414.07	A1 2004	0318 DE 200	2-10241407	20020906						
US 2004116501	A1 2004	0617 US 200	3-654365	20030903						
CA 2497374	AA 20040	0318 CA 200	3-2497374	20030905						
WO 2004022041	A2 ·2004	0318 WO 200	WO 2003-EP9895							
WO 2004022041	A3 2004	0506								
W: AE, AG, AL,	AM, AT, AU,	AZ, BA, BB, BG	G, BR, BY, BZ,	CA, CH, CN,						
		DM, DZ, EC, E								
GM, HR, HU,	ID, IL, IN,	IS, JP, KE, K	G, KP, KR, KZ,	LC, LK, LR,						

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LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
             PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
             TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                            AU 2003-271586
     AU 2003271586
                          A1
                                20040329
                                                                    20030905
     EP 1534272
                          A2
                                20050601
                                            EP 2003-753390
                                                                    20030905
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     BR 2003014031
                          Α
                                20050705
                                             BR 2003-14031
                                                                    20030905
     CN 1678307
                          Α
                                20051005
                                             CN 2003-821089
                                                                    20030905
     JP 2005539058
                          T2
                                20051222
                                             JP 2004-533499
                                                                    20030905
     ZA 2005001582
                          Α
                                20050909
                                             ZA 2005-1582
                                                                    20050222
     NO 2005001468
                          Α
                                20050603
                                             NO 2005-1468
                                                                    20050321
PRIORITY APPLN. INFO.:
                                            .DE 2002-10241407
                                                                 Α
                                                                    20020906
                                             WO 2003-EP9895
                                                                    20030905
OTHER SOURCE(S):
                         MARPAT 140:247052
     The invention discloses the use of hydroxyindolylglyoxylic acid amides as
     inhibitors of the phosphodiesterase 4 for the treatment of nonallergic
     rhinitis.
                     CAPLUS COPYRIGHT 2006 ACS on STN
L24 ANSWER 7 OF 14
ACCESSION NUMBER:
                         2004:130977 CAPLUS
DOCUMENT NUMBER:
                         140:281023
TITLE:
                         Anti-inflammatory potential of the selective
                         phosphodiesterase 4 inhibitor N-(3,5-dichloro-pyrid-4-
                         yl) - [1-(4-fluorobenzyl) -5-hydroxy-indole-3-yl] -
                         glyoxylic acid amide (AWD 12-281), in human cell
                         preparations
AUTHOR (S):
                         Draheim, Regina; Egerland, Ute; Rundfeldt,
                         Chris
CORPORATE SOURCE:
                         Departments of Pharmacology and Molecular Biology,
                         Elbion AG, Radebeul, Germany
SOURCE:
                         Journal of Pharmacology and Experimental Therapeutics
                          (2004), 308(2), 555-563
                         CODEN: JPETAB; ISSN: 0022-3565
PUBLISHER:
                         American Society for Pharmacology and Experimental
                         Therapeutics
DOCUMENT TYPE:
                         Journal
LANGUAGE:
                         English
     AWD 12-281 is a potent (IC50 = 9.7 nM) and highly selective inhibitor of
     the phosphodiesterase 4 (PDE4) isoenzyme with low affinity to the
     high-affinity rolipram-binding site. The compound was optimized for topical
     treatment of asthma, chronic obstructive pulmonary disease (COPD), and
     allergic rhinitis. The aim of the present study was to assess the effect
```

the phosphodiesterase 4 (PDE4) isoenzyme with low affinity to the high-affinity rolipram-binding site. The compound was optimized for topical treatment of asthma, chronic obstructive pulmonary disease (COPD), and allergic rhinitis. The aim of the present study was to assess the effect of AWD 12-281 in human inflammatory cells. Peripheral blood mononuclear cells (PBMCs), diluted whole blood, and human nasal polyp cells derived from surgically resected nasal polyps from patients with polyposis comprise sources of target tissue cells that can be used to predict anti-inflammatory effects in patients. AWD 12-281 was capable of suppressing the production of cytokines in stimulated PBMCs: interleukin-2 (IL-2, phytohemagglutinin stimulation), IL-5 (Con A stimulation), IL-5 and IL-4 (anti-CD3/anti-CD28 co-stimulation), and lipopolysaccharidestimulated release of tumor necrosis factor α (TNF α). The corresponding values for half-maximum inhibition, EC50, for AWD 12-281 were within a narrow range (46-121 nM). Comparing the effect of AWD 12-281 with roflumilast, cilomilast (SB 207499), rolipram (RPR-73401), and

Saloni Sharma 05/23/2006

1-(3-nitrophenyl)-3-(4-pyridylmethyl)pyrido[2,3-d]pyrimidin-2,4(1H,3H)-dione (RS-25344-000), it could be shown that the PDE4 inhibitory activity was closely correlated with inhibitory potential as measured by the above-described assays. AWD 12-281 was also shown to suppress TNF α release in dispersed nasal polyps (EC50 = 111 nM) and in diluted whole blood (EC50 = 934 nM). The reduced activity in human blood may be related to high plasma protein binding. Currently, phase II clin. studies are under way to evaluate the therapeutic potential of AWD 12-281 in asthma, COPD, and allergic rhinitis.

REFERENCE COUNT:

THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

40

ACCESSION NUMBER:

2004:60309 CAPLUS

DOCUMENT NUMBER:

140:105273

TITLE:

Topical treatment of skin diseases

INVENTOR(S):

Rundfeldt, Chris; Kietzmann, Manfred; Hoppmann, Joachim; Baeumer, Wolfgang; Kuss,

Hildegard; Hoefgen, Norbert

PATENT ASSIGNEE(S):

Elbion AG, Germany

SOURCE:

PCT Int. Appl., 48 pp.

DOCUMENT TYPE:

Patent

LANGUAGE:

English

CODEN: PIXXD2

FAMILY ACC. NUM. COUNT:

: 1

PATENT INFORMATION:

	PATENT NO.											DATE							
	WO	2004006920				A1 20040122													
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			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE	, KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,	
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							AA 20040122 CA 2003-2492093						093						
		2003							20040202 AU 2003-254332 200307										
								2005	50426 BR 2003-12696 200307							710			
	EP	1531	818			A 1		2005	0525	EP 2003-763810 2						0030	710		
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OTHER SOURCE(S): MARPAT 140:105273

AB The present invention relates to a method for the treatment of an inflammatory and/or allergic skin disease comprising topically administering a substituted hydroxy indole which is a phosphodiesterase 4 inhibitor. Examples are provided of the topical effectiveness of AWD 12-281 and cilomilast in dermal immunol. inflammation.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS

Saloni Sharma 05/23/2006

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:2852 CAPLUS

DOCUMENT NUMBER: 140:59520

TITLE: Preparation of pyrrolidine and piperidinecarboxamides as inhibitors of phosphodiesterase IV (PDE 4)

INVENTOR(S): Egerland, Ute; Rueger, Carla; Schindler, Rudolf; Rundfeldt, Chris; Kuss, Hildegard; Lichoscherstow, Arkadi M.; Seredenin, Sergey B.; Borissenko, Sergey A.

PATENT ASSIGNEE(S): Elbion A.-G., Germany SOURCE: PCT Int. Appl., 79 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO.					KIN	D :	DATE		٠, ١	APPL	ICAT:	ION 1	DATE .				
WO	WO 2004000806 .			A1 20031231			1	NO 2	003-1	EP65	20030623						
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		LS,	LT,	LU,	LV,	MA,	MD,	MG,	ΜK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,	OM,
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		TT,	ΤŹ,	UA,	ŪG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	zw				
	RW:	GH,	GM,	KΕ,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,	BY,
		KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,
		FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,
		BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG
								DE 2002-10228132									
AU 2003245979				A1 20040106 AU 2003-245979							20030623						
PRIORITY	APP	LN.	INFO	.:					1	DE 2	002-	1022	8132	1	A 2	0020	524
									1	WO 2	0.03 - 1	EP65	90	1	W 2	0030	523

OTHER SOURCE(S): MARPAT 140:59520

GI

Title compds. [I; n = 1, 2; X = NH2, N:CR3R4; NHCHR3R4; NR3CHR3R4; NHCH2R4, NHCOR4; R1, R4 = (substituted) 3-14 membered (saturated) (poly)cyclyl; 5-15 membered (saturated) (poly)heterocyclyl; R2 = H, (substituted) (branched) alkyl, PhCH2; NR1R2 = (substituted) heterocyclyl, R3 = H, (substituted) (branched) alkyl], were prepared Thus, 1-amino-pyrrolidine-2-carboxylic acid, 2,6-dichlorophenylamide, and 3,4-dimethoxybenzaldehyde in 2-propanol were refluxed for 4 h to give 84% N-(2,6-dichlorophenyl)-(E)-1-([(3,4-dimethoxyphenyl)methylene]amino)pyrrol idine-2-carboxamide. Several I at 114-5,000 nmol/L inhibited PDE 4 with IC50 = 32.4-79.6%.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Saloni Sharma

L24 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:775804 CAPLUS

DOCUMENT NUMBER: 140:104940

TITLE: In vivo efficacy in airway disease models of

N-(3,5-dichloropyrid-4-yl)-[1-(4-fluorobenzyl)-5-hydroxyindole-3-yl]glyoxylic acid amide (AWD 12-281), a selective phosphodiesterase 4 inhibitor for inhaled

administration

AUTHOR(S): Kuss, H.; Hoefgen, N.; Johanssen, S.;

Kronbach, T.; Rundfeldt, C.

CORPORATE SOURCE: Department of Pharmacology, Elbion AG, Radebeul,

Germany

SOURCE: Journal of Pharmacology and Experimental Therapeutics

(2003), 307(1), 373-385

CODEN: JPETAB; ISSN: 0022-3565

PUBLISHER: American Society for Pharmacology and Experimental

Therapeutics

DOCUMENT TYPE: Journal LANGUAGE: English

AWD 12-281 is a highly potent and selective phosphodiesterase 4 (PDE4) inhibitor that was designed to have a metabolic profile that was optimized for topical administration. The aim of the current study was to explore the pharmacol. profile of intratracheally administered AWD 12-281 in different models of asthma and chronic obstructive pulmonary disease (COPD) in comparison with steroids. To assess the anti-inflammatory potential of AWD 12-281, the antigen-induced cell infiltration in bronchoalveolar lavage fluid (BALF) of Brown Norway rats was determined AWD 12-281 (ID50 of 7 $\mu g/kg$ i.t.) as well as beclomethasone (0.1 $\mu g/kg$ i.t.) suppresses late-phase eosinophilia when administered intrapulmonary. Furthermore, AWD 12-281 has also strong anti-inflammatory properties when tested in lipopolysaccharide-induced acute lung neutrophilia in Lewis rats (ID50 of 0.02 μ g/kg i.t.), ferrets (ID50 of 10 μ g/kg i.t.), and domestic pigs (2-4 mg/pig i.t. or 1 mg/kg i.v.). In pigs, AWD 12-281 was as effective as beclomethasone (0.4 mg/pig i.t.) and dexamethasone (0.28 mg/kg i.v.), although at 3 to 10 times the dosage. The bronchodilatory activity of AWD 12-281 was assessed in sensitized guinea pigs. AWD 12-281 (1.5 mg/kg i.t., 1-h pretreatment) inhibited allergen-induced bronchoconstriction by 68% (parameter airway resistance). In sensitized BP-2 mice AWD 12-281 abolished the allergen-induced bronchial hyperresponsiveness and eosinophilia in BALF, showing dose dependence. When given orally, i.v. or i.t., AWD 12-281 has a considerably lower emetic potential than cilomilast in ferrets and roflumilast in pigs. When given topically by inhalation, no emesis could be induced in dogs up to the highest feasible dose (15 mg/kg in 50% lactose blend). These results indicate that AWD 12-281 is a unique potential new drug for the topical treatment of asthma and COPD.

REFERENCE COUNT: 43

THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:695438 CAPLUS

DOCUMENT NUMBER: 140:87294

TITLE: AWD 12-281, a highly selective phosphodiesterase 4

inhibitor, is effective in the prevention and treatment of inflammatory reactions in a model of

allergic dermatitis

AUTHOR(S): Baeumer, Wolfgang; Gorr, Gilbert; Hoppmann, Joachim;

Ehinger, Andreas M.; Rundfeldt, Chris;

Kietzmann, Manfred

CORPORATE SOURCE: Department of Pharmacology, Toxicology and Pharmacy,

School of Veterinary Medicine, Hannover, D-30559,

Germany

SOURCE: Journal of Pharmacy and Pharmacology (2003), 55(8),

1107-1114

CODEN: JPPMAB; ISSN: 0022-3573

PUBLISHER: Pharmaceutical Press

DOCUMENT TYPE: Journal LANGUAGE: English

AWD 12-281 (N-(3,5-dichloro-4-pyridinyl)-2-[1-(4-fluorobenzyl)-5-hydroxy-1H-indol-3-yl]-2-oxoacetamide), a phosphodiesterase 4 inhibitor, which is optimized for topical administration, was tested in a model of allergic dermatitis in mice. To obtain an allergic dermatitis, BALB/c mice were sensitized to toluene-2,4-diisocyanate (TDI). The allergic reaction was challenged by topical administration of TDI onto the mice ears. AWD 12-281 was tested for its anti-inflammatory potential by oral, i.p. and topical administration. The phosphodiesterase 4 inhibitor, cilomilast (SB 207499), and/or the corticosteroid, diflorasone diacetate, were used as reference compds. Given orally and i.p. 2 h before as well as 5 and 24 h after TDI challenge, AWD 12-281 showed no, or only a transient inhibition of the allergen-induced ear swelling, whereas cilomilast significantly inhibited this ear swelling. Applied topically onto the ears before TDI challenge, AWD 12-281, cilomilast and diflorasone diacetate caused total inhibition of ear swelling 24 h after challenge, confirmed by a decrease of the pro-inflammatory cytokines interleukin-4, interleukin-6 and macrophage inhibitory protein-2. Administered topically after TDI challenge as therapeutic intervention, AWD 12-281 and diflorasone diacetate caused significant inhibition of ear swelling; cilomilast failed to do so. results indicate that topically administered AWD 12-281 may be potent in the prevention and treatment of allergic/inflammatory skin diseases.

REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:420229 CAPLUS

DOCUMENT NUMBER: 138:18980 TITLE: AWD 12-281

AUTHOR(S): Kuss, H.; Hofgen, N.; Egerland,

U.; Heer, S.; Marx, D.; Szelenyi, I.; Schupke, H.; Gasparic, A.; Olbrich, M.; Hempel, R.; Hartenhauer,

H.; Krone, D.; Berthold, K.; Kronbach, T.;

Rundfeldt, C.

CORPORATE SOURCE: Arzneimittelwerk Dresden GmbH, Radebeul, D-01445,

Germany

SOURCE: Drugs of the Future (2002), 27(2), 111-116

CODEN: DRFUD4; ISSN: 0377-8282

PUBLISHER: Prous Science

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. Airway diseases such as bronchial asthma and chronic obstructive pulmonary disease (COPD) are chronic inflammatory diseases whose prevalence is increasing. Current research concerned with developing effective treatments for these conditions have focused on the search for alternatives to the standard corticosteroid antiinflammatory therapy. Selective phosphodiesterase 4 (PDE4) inhibitors have received a considerable amount of attention due to their ability to suppress the functions of several cell types involved in allergic and inflammatory disorders. The selective PDE4 inhibitor AWD 12-281 is the result of a

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pharmacophore-based synthesis program wherein the optimization process was supported by ligand-based drug design methods. AWD 12-281 was selected for further development for its high affinity and selectivity for the human PDE4 isoenzyme and due to its potent activity and excellent tolerability in models of allergic rhinitis, asthma and COPD, especially after topical treatment.

REFERENCE COUNT:

39 THERE ARE 39 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2000:55462 CAPLUS

DOCUMENT NUMBER:

132:202635

TITLE:

A peptidic binding site model for PDE 4 inhibitors

AUTHOR (S):

CORPORATE SOURCE:

Polymeropoulos, Emmanuel E.; Hofgen, Norbert Department of Chemical Research, Corporate R and D

ASTA Medica Group, Frankfurt, D-60314, Germany

SOURCE:

Quantitative Structure-Activity Relationships (1999),

18(6), 543-547

CODEN: QSARDI; ISSN: 0931-8771

PUBLISHER:

Wiley-VCH Verlag GmbH.

DOCUMENT TYPE:

Journal

LANGUAGE:

English

The pseudoreceptor modeling program PrGen was used to construct a peptidic binding site model for phosphodiesterase 4 inhibitors. A training set of 21 diverse compds. (rolipram, nitraquazone and xanthine derivs., imidazo pyrido pyrazinones and 5-oxyindoles) was used to construct the binding site surrogate consisting of five amino acid residues, a Zn+2 cofactor and an envelope of charged virtual particles. The model was validated by predicting the free energies of binding AGpred0 of ten ligands (rolipram, imidazo pyrido pyrazinones and 5-oxyindoles). In seven cases the prediction was satisfactory. The rms deviation [4] in $\Delta G0$ is 0.16 and 1.82 kcal/mol-resulting in an uncertainty in IC50 (or Ki) of 1.32 and 22.81-for the training and the test set resp., while the corresponding maximal prediction errors in ΔGpred0 were 0.27 kcal/mol and 4.50 kcal/mol.

REFERENCE COUNT:

15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1999:708761 CAPLUS

DOCUMENT NUMBER:

131:310549

TITLE:

New hydroxyindoles and their use as phosphodiesterase

4 and TNF α inhibitors

INVENTOR (S):

Hofgen, Norbert; Egerland, Ute; Poppe,

Hildegard; Marx, Degenhard; Szelenyi, Stefan; Kronbach, Thomas; Polymeropoulos, Emmanuel; Heer,

Sabine

PATENT ASSIGNEE(S):

Arzneimittelwerk Dresden GmbH, Germany

PCT Int. Appl., 45 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

. Patent

LANGUAGE:

SOURCE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
WO 9955696	A1 (19991104	WO 1999-EP2792	19990424
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                                              DE 1998-19818964
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                                              WO 1999-EP2792
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                                              US 2002-81642
                                                                  A1 20020221
                                              US 2002-81807
                                                                  A3 20020221
                          MARPAT 131:310549
 OTHER SOURCE(S):
GI
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Saloni Sharma 05/23/20'

AB Hydroxyindoles I [R1, R5 = (un)substituted aliphatic, carbocyclic, heterocyclic, spirocyclic; R2, R3 = H, OH, ≥1 of them being OH; R4 = H, (un)substituted OH, SH, S(O)H, SO2H, NH2, CO2H, C(S)OH, NO2, CN, F, Cl, Br, I; A = alkylene, alkenylene, (CHOZ)m, CO, CS, C:NZ, O, S, NZ; Z = (un)substituted alkyl, alkenyl, carbocyclic, heterocyclic; B = C, S, SO; D = O, S, CH2, NZ; E = bond, (CH2)m, O, S, NZ; m = 0-3] were prepared I have IC50 for PDE IV inhibition of 1X10-9-1X10-5 and a selectivity relative to PDE's 2, 3, and 5 of 100-10,000. N-(3,5-dichloro-4-pyridyl)-2-[1-(4-fluorobenzyl)-5-methoxy-3-indolyl]-2-oxoacatamide was obtained by demethylation of the 5-methoxy compound and was reduced to the 2-hydroxyacetamide with NaBH4.

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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NEW FILES RELEASED
***Regulatory Affairs Journals (File 183)
***Index Chemicus (File 302)
***Inspec (File 202)
RESUMED UPDATING
***File 141, Reader's Guide Abstracts
RELOADS COMPLETED
***File 516, D&B--Dun's Market Identifiers
***File 523, D&B European Dun's Market Identifiers
***File 531, American Business Directory
*** MEDLINE has been reloaded with the 2006 MeSH (Files 154 & 155)
*** The 2005 reload of the CLAIMS files (Files 340, 341, 942)
is now available online.
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DATABASES REMOVED
***File 196, FINDEX
***File 468, Public Opinion Online (POLL)
Chemical Structure Searching now available in Prous Science Drug
Data Report (F452), Prous Science Drugs of the Future (F453),
IMS R&D Focus (F445/955), Pharmaprojects (F128/928), Beilstein
Facts (F390), Derwent Chemistry Resource (F355) and Index Chemicus
(File 302).
                   ***
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 >>>and events, please visit What's New from Dialog at <<<
 >>>http://www.dialog.com/whatsnew/. You can find news about<<<
 >>>a specific database by entering HELP NEWS <file number>.<<
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(c) 2006 BIOSIS. All rts. reserv.
0014007816
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Preclinical evidence of azelastine hydrochloride activity
AUTHOR: Lieberman Phillip (Reprint)
AUTHOR ADDRESS: 300 Walnut Bend Road South, Cordova, TN, 38018, USA**USA
JOURNAL: Current Therapeutic Research 63 (9): p556-571 September, 2002
2002
MEDIUM: print
ISSN: 0011-393X
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
 2002
... ABSTRACT: literature published in the previous 15 years. The search
  terms used were azelastine, allergic rhinitis, nonallergic rhinitis,
  and anti-inflammatory activity. Results: In addition to having
 H1-blocking activity, azelastine has broad-based anti-inflammatory
  activity, probably accounting for its utility in the treatment of
               rhinitis . The mechanisms governing its anti-inflammatory
  activity include prevention of mast cell and basophil degranulation...
DESCRIPTORS:
  ...MAJOR CONCEPTS: Respiratory System...
... Respiration
  ...DISEASES: immune system disease, respiratory system disease, drug
    therapy...
... nonallergic rhinitis --...
... respiratory system disease, drug therapy
5/3, K/2
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(c) 2006 BIOSIS. All rts. reserv.
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Review of the upper airway, including olfaction, as mediator of symptoms
AUTHOR: Shusterman Dennis (Reprint)
AUTHOR ADDRESS: Upper Airway Biology Laboratory, University of California,
  1301 S. 46th St., Bldg. 112, Richmond, CA, 94804, USA**USA
JOURNAL: Environmental Health Perspectives 110 (Supplement 4): p649-653
August, 2002 2002
MEDIUM: print
ISSN: 0091-6765
DOCUMENT TYPE: Article; Literature Review
RECORD TYPE: Abstract
LANGUAGE: English
 2002
... ABSTRACT: the chemical qualities of the air we breathe. A number of
 poorly understood conditions, including nonallergic rhinitis,
  irritant-induced rhinitis, odor-triggered asthma, odor-triggered panic
  attacks, chemical-induced olfactory dysfunction, and...
DESCRIPTORS:
  ...ORGANISMS: PARTS ETC: respiratory system
  ...DISEASES: respiratory system disease...
... respiratory system disease...
... respiratory system disease...
... nonallergic rhinitis --...
... respiratory system disease...
...immune system disease, respiratory system disease
 5/3, K/3
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0013846124
            BIOSIS NO.: 200200439635
Immunohistochemical localization of subtypes of muscarinic receptors in
 human inferior turbinate mucosa
AUTHOR: Nakaya Muneo (Reprint); Yuasa Takafumi; Usui Nobuo
AUTHOR ADDRESS: Dept of Otolaryngology, Tokyo University, Hongo 7-3-1,
 Bunkyo-Ku, 113-8655, Tokyo, Japan**Japan
JOURNAL: Annals of Otology Rhinology and Laryngology 111 (7): p593-597
July, 2002 2002
MEDIUM: print
ISSN: 0003-4894
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
2002
... ABSTRACT: the best target for more selective muscarinic drugs and quide
 the treatment of allergic and nonallergic
DESCRIPTORS:
 MAJOR CONCEPTS: Respiratory System...
... Respiration
```

...ORGANISMS: PARTS ETC: respiratory system

```
5/3,K/4
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 200200384741
0013791230
Fluticasone propionate downregulates nasal fibroblast functions involved in
  airway inflammation and remodeling
AUTHOR: Silvestri M; Sabatini F; Scarso L; Cordone A; Dasic G; Rossi G A
  (Reprint)
AUTHOR ADDRESS: Divisione di Pneumologia, Istituto G. Gaslini, Largo G.
  Gaslini, 5, I-16148, Genoa, Italy**Italy
JOURNAL: International Archives of Allergy and Immunology 128 (1): p51-58
May, 2002 2002
MEDIUM: print
ISSN: 1018-2438
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
 2002
ABSTRACT: Background: Besides being highly effective in the treatment of
  allergic and nonallergic rhinitis with eosinophilia, intranasal
  corticosteroids appear to be useful in reducing nasal polypoid lesions
  and the...
DESCRIPTORS:
  ...ORGANISMS: PARTS ETC: respiratory system, inflammation, remodeling
... respiratory system, function, regulation
  ...DISEASES: immune system disease, respiratory system disease
5/3, K/5
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0013678385
            BIOSIS NO.: 200200271896
Nasal lavage concentrations of free hemoglobin as a marker of
 microepistaxis during nasal provocation testing
AUTHOR: Park Y-J; Repka-Ramirez M S; Naranch K; Velarde A; Clauw D;
  Baraniuk J N (Reprint)
AUTHOR ADDRESS: Division of Rheumatology, Immunology and Allergy,
  Georgetown University, 3800 Reservoir Road, LL Gorman Building,
  Georgetown, USA**USA
JOURNAL: Allergy (Copenhagen) 57 (4): p329-335 April, 2002 2002
MEDIUM: print
ISSN: 0105-4538
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
2002
... ABSTRACT: saline nasal provocation. Unilateral hypertonic nasal
 provocation was performed in normal, allergic rhinitis (AR) and
               rhinitis (NAR) subjects (total of 1316 specimens). fHb was
 nonallergic
 measured using the Sigma-Aldrich kit (St...
```

...16.5 mug/ml. Elevations of fHb without changes in albumin were more

prevalent in nonallergic rhinitis . Conclusions: Significant bleeding into nasal lavage samples can contaminate the specimens and increase the concentrations... **DESCRIPTORS:** ...DISEASES: immune system disease, respiratory system disease... ... nonallergic rhinitis --... ... respiratory system disease CHEMICALS & BIOCHEMICALS: ...allergic rhinitis study, microepistaxis marker, nasal lavage fluid concentration, nasal provocation testing effects, nonallergic rhinitis study 5/3,K/6 DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv. 0013098525 BIOSIS NO.: 200100270364 Intranasal beclomethasone dipropionate in the treatment of common cold AUTHOR: Qvarnberg Yrjo (Reprint); Valtonen Hannu; Laurikainen Kari AUTHOR ADDRESS: Department of Otorhinolaryngology, Central Hospital of Central Finland, FIN-40620, Jyvaskyla, Finland**Finland JOURNAL: Rhinology (Utrecht) 39 (1): p9-12 March, 2001 2001 MEDIUM: print ISSN: 0300-0729 DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English 2001 ABSTRACT: Sinusitis is usually considered a complication of viral rhinitis. Virus infections in the upper respiratory tract lead to mucosal swelling, which may obstruct paranasal sinus outflow, resulting in infection in... ...been found beneficial in a variety of acute and chronic nasal conditions including allergic and nonallergic rhinitis and chronic rhinosinusitis. The purpose of this study was to examine whether the intranasal inhalation... DESCRIPTORS: ...ORGANISMS: PARTS ETC: respiratory system... ...upper respiratory tract... ... respiratory system ...DISEASES: immune system disease, respiratory system disease... ... respiratory system disease, viral disease... ... respiratory system disease... ... respiratory system disease... ... respiratory system disease...

... respiratory system disease, viral disease

```
5/3,K/7
DIALOG(R)File 5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0013002011
            BIOSIS NO.: 200100173850
Expression of costimulatory CD80/CD86-CD28/CD152 molecules in nasal mucosa
  of patients with perennial allergic rhinitis
AUTHOR: Okano Mitsuhiro (Reprint); Hattori Hisashi (Reprint); Nagano
  Toshiaki; Takishita Teruaki (Reprint); Azuma Miyuki; Nishizaki Kazunori
  (Reprint)
AUTHOR ADDRESS: Okayama University Medical School, Okayama, Japan**Japan
JOURNAL: Journal of Allergy and Clinical Immunology 107 (2): pS150
February, 2001 2001
MEDIUM: print
CONFERENCE/MEETING: 57th Annual Meeting of the American Academy of Allergy,
Asthma and Immunology New Orleans, Louisiana, USA March 16-21, 2001;
20010316
SPONSOR: American Academy of Allergy Asthma and Immunology
ISSN: 0091-6749
DOCUMENT TYPE: Meeting; Meeting Abstract
RECORD TYPE: Citation
LANGUAGE: English
 2001
DESCRIPTORS:
  ...ORGANISMS: PARTS ETC: respiratory system
  ...DISEASES: immune system disease, respiratory system disease,
    perennial...
... nonallergic rhinitis --...
... respiratory system disease
 5/3,K/8
DIALOG(R) File
              5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
           BIOSIS NO.: 200100145241
0012973402
Inflammatory cell and epithelial characteristics of perennial allergic and
              rhinitis with a symptom history of 1 to 3 years' duration
  nonallergic
AUTHOR: Amin Kawa (Reprint); Rinne Juhani; Haahtela Tari; Simola Markku;
  Peterson Christer G B; Roomans Godfried M; Malmberg Henrik; Venge Per;
  Seveus Lahja
AUTHOR ADDRESS: Department of Genetics and Pathology, Rudbecks Laboratory,
  University of Uppsala, SE-751 85, Uppsala, Sweden**Sweden
JOURNAL: Journal of Allergy and Clinical Immunology 107 (2 Part 2): p
249-257 February, 2001 2001
MEDIUM: print
ISSN: 0091-6749
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
Inflammatory cell and epithelial characteristics of perennial allergic and
 nonallergic rhinitis with a symptom history of 1 to 3 years' duration
 2001
```

... ABSTRACT: were obtained from 27 patients with perennial allergic

rhinitis (PAR), from 12 patients with perennial nonallergic rhinitis

```
(PNAR) with eosinophils present in the nasal smear, and from 6 control
  subjects without rhinitis ...
DESCRIPTORS:
  ...DISEASES: respiratory system disease, 1 to 3 year symptom history,
    allergic type, epithelial characteristics, inflammatory cell
    characteristics
 5/3,K/9
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 200000383967
0012665654
Binding of glucocorticoids to human nasal tissue in vitro
AUTHOR: Esmailpour Nasser; Hoegger Petra; Rohdewald Peter (Reprint)
AUTHOR ADDRESS: Institute of Pharmaceutical Chemistry, Westfaelische
  Wilhelms-Universitaet, Hittorfstrasse 58-62, D-48149, Muenster, Germany**
  Germany
JOURNAL: International Archives of Allergy and Immunology 122 (2): p
151-154 June, 2000 2000
MEDIUM: print
ISSN: 1018-2438
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
 2000
... ABSTRACT: Intranasal application of glucocorticoids is an efficacious
 treatment of allergic rhinitis and some cases of nonallergic rhinitis
  . However, no data on binding of glucocorticoids to nasal tissue are
  available. Pronounced binding of ...
DESCRIPTORS:
  ... MAJOR CONCEPTS: Respiratory System...
... Respiration
  ...ORGANISMS: PARTS ETC: respiratory system...
... respiratory system disease
  ...DISEASES: immune system disease, respiratory system disease...
... nonallergic
                 rhinitis --...
...immune system disease, respiratory system disease
 5/3,K/10
DIALOG(R)File
                5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0012661216
            BIOSIS NO.: 200000379529
Partial choanal atresia masking as allergic rhinitis
AUTHOR: Cox Ronald L (Reprint); Freeman Theodore M
AUTHOR ADDRESS: Department of Allergy and Immunology, Wilford Hall Medical
 Center, United States Air Force, 59th Medical Wing Lackland Air Force
  Base, San Antonio, TX, 78236, USA**USA
JOURNAL: Pediatric Asthma Allergy and Immunology 14 (2): p129-135 Summer,
2000 2000
MEDIUM: print
```

ISSN: 0883-1874

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

2000

...ABSTRACT: of approximately 30% of the population. Rhinitis can be classified into allergic rhinitis (AR) and **nonallergic rhinitis** (NAR). An interesting case of NAR is presented of a 4-year-old female with...

DESCRIPTORS:

- ...DISEASES: immune system disease, respiratory system disease...
- ...congenital disease, respiratory system disease

5/3,K/11

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0012624493 BIOSIS NO.: 200000342806

Immunolocalization of inducible nitric oxide synthase and 3-nitrotyrosine in the nasal mucosa of patients with rhinitis

AUTHOR: Kang Bor-Hwang; Chen Shinn-Shong; Jou Lin-Shu; Weng Pinh-Kun; Wang Hsing-Won (Reprint)

AUTHOR ADDRESS: Department of Otolaryngology, Tri-Service General Hospital, Ting-Chow Rd, 8 Section 3, Taipei, 100, Taiwan**Taiwan

JOURNAL: European Archives of Oto-Rhino-Laryngology 257 (5): p242-246 May, 2000 2000

MEDIUM: print ISSN: 0937-4477

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

2000

- ...ABSTRACT: peroxynitrite in the pathogenesis of rhinitis. Inferior nasal turbinates were obtained from allergic rhinitis and **nonallergic rhinitis** patients during corrective nasal surgery. The expressions of the inducible form of nitric oxide synthase...

 DESCRIPTORS:
 - ...MAJOR CONCEPTS: Respiratory System...

... Respiration

...ORGANISMS: PARTS ETC: respiratory system ...DISEASES: respiratory system disease

5/3,K/12

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0012360297 BIOSIS NO.: 200000078610

Medical management of sinusitis

AUTHOR: Kaliner Michael (Reprint)

AUTHOR ADDRESS: Institute for Asthma and Allergy, Washington Hospital Center, Washington, DC, USA**USA

JOURNAL: American Journal of the Medical Sciences 316 (1): p21-28 July, 1998 1998

MEDIUM: print ISSN: 0002-9629

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

1998

...ABSTRACT: abnormalities, and sinonasal microbiology. The most common events leading to sinusitis are colds, allergic and **nonallergic rhinitis**, and anatomic defects which interfere with the sinus outflow tracks. Treatment involves drainage of the...

DESCRIPTORS:

...ORGANISMS: PARTS ETC: respiratory system, drainage ...DISEASES: respiratory system disease, management

5/3,K/13

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0012217795 BIOSIS NO.: 199900477455

A controlled study on the effectiveness of loratadine in combination with flunisolide in the treatment of nonallergic rhinitis with eosinophilia (NARES)

AUTHOR: Purello-D'Ambrosio F (Reprint); Isola S; Ricciardi L; Gangemi S; Barresi L; Bagnato G F

AUTHOR ADDRESS: Istituto di Patologia Medica, Policlinico Universitario, Padiglione H, 98122, Messina, Italy**Italy

JOURNAL: Clinical and Experimental Allergy 29 (8): p1143-1147 Aug., 1999 1999

MEDIUM: print ISSN: 0954-7894

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

...controlled study on the effectiveness of loratadine in combination with flunisolide in the treatment of nonallergic rhinitis with eosinophilia (NARES)

1999

ABSTRACT: Background Nonallergic rhinitis with eosinophilia (NARES), accounting for some 15% of perennial rhinitis, is a nasal disorder whose

DESCRIPTORS:

DISEASES: nonallergic rhinitis with eosinophilia...

...blood and lymphatic disease, respiratory system disease

5/3,K/14

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0012071140 BIOSIS NO.: 199900330800

Serum-soluble Fas levels as a marker to distinguish allergic and nonallergic rhinitis

AUTHOR: Kato Masashi (Reprint); Hattori Taku; Ito Hirotaka; Kageyama Motoo; Yamashita Tetsuji; Nitta Yukiko; Nakashima Izumi

AUTHOR ADDRESS: Department of Immunology, Nagoya University School of

```
Medicine, 65 Tsurumai-cho, Showa-ku, Nagoya, Aichi, 466-8550, Japan**
  Japan
JOURNAL: Journal of Allergy and Clinical Immunology 103 (6): p1213-1214
June, 1999 1999
MEDIUM: print
ISSN: 0091-6749
DOCUMENT TYPE: Article
RECORD TYPE: Citation
LANGUAGE: English
Serum-soluble Fas levels as a marker to distinguish allergic and
  nonallergic rhinitis
 1999
DESCRIPTORS:
  ...DISEASES: immune system disease, respiratory system disease...
... nonallergic
                 rhinitis --...
... respiratory system disease
 5/3,K/15
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0011895478
            BIOSIS NO.: 199900155138
ICAM-1 expression on sputum cells as a marker of persistent inflammation in
  patients with allergic and nonallergic rhinitis and in asthmatics
AUTHOR: Foresi A (Reprint); Leone C; Teodoro C; Mastropasqua B; Pelucchi A;
  Chetta A; Fichera E; Burlone E; Olivieri D
AUTHOR ADDRESS: Serv. Fisopat. Respir., Sesto S. Giovanni, Parma, Italy**
  Italv
JOURNAL: European Respiratory Journal 12 (SUPPL. 28): p370S Sept., 1998
1998
MEDIUM: print
CONFERENCE/MEETING: European Respiratory Society Annual Congress Geneva,
Switzerland September 19-23, 1998; 19980919
SPONSOR: The European Respiratory Society
ISSN: 0903-1936
DOCUMENT TYPE: Meeting; Meeting Abstract
RECORD TYPE: Citation
LANGUAGE: English
... expression on sputum cells as a marker of persistent inflammation in
 patients with allergic and nonallergic rhinitis and in asthmatics
 1998
DESCRIPTORS:
  ... MAJOR CONCEPTS: Respiratory System...
... Respiration
  ...ORGANISMS: PARTS ETC: respiratory system
  ...DISEASES: immune system disease, respiratory system disease...
...immune system disease, respiratory system disease...
... respiratory system disease
 5/3, K/16
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
```

0011829707 BIOSIS NO.: 199900089367

Parameters for the diagnosis and management of sinusitis

AUTHOR: Joint Task Force On Practice Parameters

JOURNAL: Journal of Allergy and Clinical Immunology 102 (6 PART 2): p

1A-9A, S107-S144 Dec., 1998 1998

MEDIUM: print ISSN: 0091-6749

DOCUMENT TYPE: Article; Standard

RECORD TYPE: Abstract LANGUAGE: English

1998

- ...ABSTRACT: as "acute" when lasting 3 to 8 weeks and "chronic" when lasting longer. Viral upper respiratory infections frequently precede subsequent bacterial invasion of the sinuses by Streptococcus pneumoniae, Haemophilus influenzae, and...
- ...tests for immunodeficiency. Nasal cytology is useful in the clinical evaluation of underlying allergic rhinitis, nonallergic rhinitis with eosinophilia syndrome, nasal polyposis, and aspirin-sensitive patients. Quantitative sweat chloride tests for diagnosis...

 DESCRIPTORS:
 - ...DISEASES: respiratory system disease, diagnosis, practice parameters, management

5/3,K/17

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0011410854 BIOSIS NO.: 199800205101

Expression of histamine receptors in nasal epithelial cells and endothelial cells- the effects of sex hormones

AUTHOR: Hamano Nanako (Reprint); Terada Nobuhisa; Maesako Ken-Ichi; Ikeda Tatehiko; Fukuda Setsuya; Waita Jun; Yamashita Tetsuji; Konno Akiyoshi AUTHOR ADDRESS: Dep. Otorhinolaryngology, Chiba Univ. Sch. Med., 1-8-1 Inohana, Chiba 260, Japan**Japan

JOURNAL: International Archives of Allergy and Immunology 115 (3): p 220-227 March, 1998 1998

MEDIUM: print

MEDIUM: print ISSN: 1018-2438

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

1998

...ABSTRACT: We compared the expressions on the specimens from patients with nasal allergy with those with **nonallergic rhinitis** or those from normal volunteers. In addition, we investigated the effects of female hormones on...

DESCRIPTORS:

- ...ORGANISMS: PARTS ETC: respiratory system...
- ... respiratory system
 - ...DISEASES: immune system disease, respiratory system disease

```
5/3,K/18
DIALOG(R) File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199800187770
0011393523
Sense of smell in allergic and nonallergic
                                             rhinitis
AUTHOR: Simola M (Reprint); Malmberg H
AUTHOR ADDRESS: Helsinki Univ. Central Hosp., Dep. Otorhinolaryngol.,
  Haartmaninkatu 4E, 00290 Helsinki, Finland**Finland
JOURNAL: Allergy (Copenhagen) 53 (2): p190-194 Feb., 1998 1998
MEDIUM: print
ISSN: 0105-4538
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
Sense of smell in allergic and nonallergic
                                             rhinitis
1998
ABSTRACT: Hyposmia is a fairly common complaint in patients with
  long-continuing allergic or nonallergic rhinitis. Other factors such
  as aging, smoking, or nasal surgery may affect olfaction, but these have
DESCRIPTORS:
  ...DISEASES: respiratory system disease...
... nonallergic rhinitis --...
... respiratory system disease
 5/3,K/19
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
           BIOSIS NO.: 199800154447
0011360200
Quality of life of patients with allergic and nonallergic rhinitis: The
  impact of allergy consultation and caregiver
AUTHOR: Harvey R; Cvietusa P; Sanders B; Westley R; Marsh W; Williams J;
  Conner D; Beck A; Speicher B
AUTHOR ADDRESS: Kaiser Permanente, Denver, CO, USA**USA
JOURNAL: Journal of Allergy and Clinical Immunology 101 (1 PART 2): pS180
Jan., 1998 1998
MEDIUM: print
CONFERENCE/MEETING: 54th Annual Meeting of the American Academy of Allergy,
Asthma and Immunology Washington, DC, USA March 13-18, 1998; 19980313
SPONSOR: American Academy of Allergy, Asthma, and Immunology
ISSN: 0091-6749
DOCUMENT TYPE: Meeting; Meeting Abstract
RECORD TYPE: Citation
LANGUAGE: English
Quality of life of patients with allergic and nonallergic rhinitis: The
  impact of allergy consultation and caregiver
 1998
DESCRIPTORS:
  ...DISEASES: respiratory system disease, caregiver impact, treatment,
    consultation impact...
```

... respiratory system disease, caregiver impact, consultation impact

```
5/3,K/20
DIALOG(R)File
               5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
           BIOSIS NO.: 199800154176
0011359929
Nasal lavage and capillary suction obtain different compartments of the
  nasal mucociliary system
AUTHOR: Ostertag P; Rasp G; Kramer M
AUTHOR ADDRESS: ENT Univ., Munich, Germany**Germany
JOURNAL: Journal of Allergy and Clinical Immunology 101 (1 PART 2): pS113
Jan., 1998 1998
MEDIUM: print
CONFERENCE/MEETING: 54th Annual Meeting of the American Academy of Allergy,
Asthma and Immunology Washington, DC, USA March 13-18, 1998; 19980313
SPONSOR: American Academy of Allergy, Asthma, and Immunology
ISSN: 0091-6749
DOCUMENT TYPE: Meeting; Meeting Abstract
RECORD TYPE: Citation
LANGUAGE: English
1998
DESCRIPTORS:
  ...ORGANISMS: PARTS ETC: respiratory system
  ...DISEASES: immune system disease, respiratory system disease...
... nonallergic rhinitis --...
...immune system disease, respiratory system disease
 5/3,K/21
               5:Biosis Previews(R)
DIALOG(R)File
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199800115298
0011321051
Comparison of nasal mucosal responsiveness to neuronal stimulation in
  non-allergic and allergic rhinitis: Effects of capsaicin nasal challenge
AUTHOR: Sanico A M; Philip G; Proud D; Naclerio R M; Togias A (Reprint)
AUTHOR ADDRESS: Johns Hopkins Asthma Allergy Cent., Unit Office 7, 5501
 Hopkins Bayview Circle, Baltimore, MD 21224-6801, USA**USA
JOURNAL: Clinical and Experimental Allergy 28 (1): p92-100 Jan., 1998
1998
MEDIUM: print
ISSN: 0954-7894
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
 1998
... ABSTRACT: increase in albumin levels and a trend in total protein
  levels. Conclusions. We conclude that nonallergic rhinitis is not
  characterized by increased responsiveness of capsaicin-sensitive nerve
  fibres; while allergic rhinitis is...
  ...DISEASES: immune system disease, respiratory system disease...
```

... respiratory system disease

```
5/3,K/22
DIALOG(R) File
                5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199800100639
0011306392
Mucoglycoprotein hypersecretion in allergic rhinitis and cystic fibrosis
AUTHOR: Yuta Atsushi; Ali Mushtaq; Sabol Marybeth; Gaumond Ethan; Baraniuk
  James N (Reprint)
AUTHOR ADDRESS: Div. Rheumatol. Immunol. Allergy, GL-008, Gorman Building,
  Georgetown Univ. Med. Cent., 3800 Reservoir Rd. NW, Washington, DC
  20007-2197, USA**USA
JOURNAL: American Journal of Physiology 273 (6 PART 1): pL1203-L1207 Dec.,
1997 1997
MEDIUM: print
ISSN: 0002-9513
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
 1997
... ABSTRACT: information about specific changes in submucosal gland
  exocytosis in diseases such as allergic rhinitis (AR), nonallergic
           (NAR), and cystic fibrosis (CF). Nasal lavage fluids were
  collected from normal, AR, NAR, and...
DESCRIPTORS:
 MAJOR CONCEPTS: Respiratory System...
... Respiration
  ...DISEASES: respiratory system disease...
...digestive system disease, genetic disease, metabolic disease,
    respiratory system disease...
... nonallergic
                rhinitis --...
... respiratory system disease
 5/3,K/23
DIALOG(R)File
                5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199800100620
0011306373
Bronchial responsiveness and airway inflammation in patients with
              rhinitis with eosinophilia syndrome
 nonallergic
AUTHOR: Leone Clementina; Teodoro Concetta; Pelucchi Andrea; Mastropasqua
 Berardino; Cavigioli Giampaolo; Marazzini Luigi; Foresi Antonio (Reprint)
AUTHOR ADDRESS: Servizio di Fisiopatol. Respir., Viale Matteotti 83, 20099
  Sesto San Giovanni, Italy**Italy
JOURNAL: Journal of Allergy and Clinical Immunology 100 (6 PART 1): p
775-780 Dec., 1997 1997
MEDIUM: print
ISSN: 0091-6749
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
```

Bronchial responsiveness and airway inflammation in patients with

nonallergic rhinitis with eosinophilia syndrome 1997

- ABSTRACT: Background: **Nonallergic rhinitis** with eosinophilia syndrome (NARES) is characterized by persistent nasal symptoms without allergy and by a...
- ...excluded by skin prick tests and RASTs. None of the patients had a history of **respiratory** symptoms. We preliminarily performed nasal lavage in all patients, and the diagnosis of NARES was...
- ...0.001). Conclusion: We showed that 46% of patients with NARES but without histories of **respiratory** symptoms had a measurable bronchial responsiveness. The presence of bronchial responsiveness was associated with an...

DESCRIPTORS:

- ...DISEASES: respiratory system disease...
- ... nonallergic rhinitis --...
- ... respiratory system disease

5/3,K/24

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0011066129 BIOSIS NO.: 199799700189

Air pollution in relation to allergic and nonallergic rhinitis
AUTHOR: Samir Magdy (Reprint); Magdy Sabry; El Fetoh Aisha A
AUTHOR ADDRESS: 12Ibn El Wardy St., Hegaz Square, Heliopolis, Cairo, Egypt
**Egypt

JOURNAL: Archives of Otolaryngology Head and Neck Surgery 123 (7): p 746-748 1997 1997

ISSN: 0886-4470

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

Air pollution in relation to allergic and nonallergic rhinitis 1997

- ABSTRACT: Objective: To investigate the relationship between allergic and nonallergic rhinitis and the exposure to air pollution. Design: Blood cadmium levels were measured in 30 patients with allergic rhinitis, 30 patients with nonallergic rhinitis, and 16 normal control subjects using atomic absorption spectrophotometry. The cadmium level was used as
- ...Results: Blood cadmium levels were significantly high in the allergic rhinitis group compared with the **nonallergic rhinitis** and control groups (P lt .001). The mean blood cadmium level in the **nonallergic rhinitis** group was higher than that in the controls, yet the difference was statistically insignificant. Also...
- ...between air pollution and this condition. The exact mechanism, however, remains to be determined. In **nonallergic rhinitis**, it seems that the contribution of air pollution as a predisposing factor is small compared

DESCRIPTORS:

... MAJOR CONCEPTS: Respiratory System...

```
... Respiration ;
  MISCELLANEOUS TERMS:
                         ... NONALLERGIC
                                           RHINITIS ; ...
... RESPIRATORY SYSTEM...
... RESPIRATORY SYSTEM DISEASE
 5/3,K/25
DIALOG(R) File
                5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199799638416
0011004356
Rhinopharyngoscopy, computed tomography and magnetic resonance imaging
AUTHOR: Bonifazi F (Reprint); Bilo M B; Antonicelli L; Bonetti M G
AUTHOR ADDRESS: Allergy Respiratory Unit, Regional Hosp., Largo Cappelli 1,
  60100 Ancona, Italy**Italy
JOURNAL: Allergy (Copenhagen) 52 (SUPPL. 33): p28-31 1997 1997
ISSN: 0105-4538
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
 1997
... ABSTRACT: one of the most useful diagnostic tools in the complex
  differential diagnosis between allergic and nonallergic rhinitis
  Furthermore, chronic allergic rhinitis, with secondary impairment of
  mucociliary clearance and the plethora of frequent...
DESCRIPTORS:
                        ... RESPIRATORY SYSTEM...
 MISCELLANEOUS TERMS:
... RESPIRATORY SYSTEM DISEASE
 5/3,K/26
DIALOG(R)File
              5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199799621720
0010987660
Nasal polyposis, sinusitis, and nonallergic
BOOK TITLE: Allergic diseases: Diagnosis and management
AUTHOR: Bernstein David I
BOOK AUTHOR/EDITOR: Patterson R (Editor)
AUTHOR ADDRESS: Div. Immunol., ML563, Univ. Cincinnati Coll. Med.,
  Cincinnati, OH 45267-0563, USA**USA
p425-437 1997
BOOK PUBLISHER: Lippincott-Raven Publishers {a}, 227 East Washington
                  Square, Philadelphia, Pennsylvania 19106, USA
ISBN: 0-397-51609-6
DOCUMENT TYPE: Book Chapter
RECORD TYPE: Citation
LANGUAGE: English
Nasal polyposis, sinusitis, and nonallergic
                                               rhinitis
 1997
DESCRIPTORS:
  MISCELLANEOUS TERMS: ... NONALLERGIC
                                           RHINITIS ; ...
```

... RESPIRATORY SYSTEM DISEASE

5/3,K/27 5:Biosis Previews(R) DIALOG(R)File (c) 2006 BIOSIS. All rts. reserv. 0010915730 BIOSIS NO.: 199799549790 Nasal biopsy is superior to nasal smear for finding eosinophils in nonallergic rhinitis AUTHOR: Ingels K (Reprint); Durdurez J-P; Cuvelier C; Van Cauwenberge P AUTHOR ADDRESS: Academic Hosp. Nijmegen St. Radboud, Dep. Otorhinolaryngol., Geert Grooteplein Zuid 18, 6525 GA Nijmegen, Netherlands**Netherlands JOURNAL: Allergy (Copenhagen) 52 (3): p338-341 1997 1997 ISSN: 0105-4538 DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English Nasal biopsy is superior to nasal smear for finding eosinophils in nonallergic rhinitis 1997 ABSTRACT: The presence of eosinophils was compared in nasal biopsy and smear. Thirty-two nonallergic rhinitis patients, of whom six had nasal polyps, were included in the study. The specimens were... ...at least four eosinophils in four fields as hypereosinophilic, our group of patients contained 25% nonallergic rhinitis with eosinophilia syndrome (NARES) patients. DESCRIPTORS: ...MAJOR CONCEPTS: Respiratory System... ... Respiration ; MISCELLANEOUS TERMS: ... NONALLERGIC RHINITIS ; RESPIRATORY SYSTEM DISEASE 5/3,K/28 DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv. 0010810453 BIOSIS NO.: 199799444513 Patient-rated overall treatment satisfaction and effectiveness with three dosing regimens of intranasal fluticasone propionate (FP) in perennial nonallergic rhinitis (PNAR) AUTHOR: Pepsin P J (Reprint); Howlandi W C Ii; Finn A F Jr; Cox F M; Bowers B W; Montgomery E; Westlung R AUTHOR ADDRESS: HealthQuest Therapy Res. Inst. Inc., Austin, TX, USA**USA JOURNAL: Journal of Allergy and Clinical Immunology 99 (1 PART 2): pS442 CONFERENCE/MEETING: Joint Meeting of the American Academy of Allergy, Asthma and Immunology, the American Association of Immunologists and the Clinical Immunology Society San Francisco, California, USA February 21-26, 1997; 19970221 ISSN: 0091-6749 DOCUMENT TYPE: Meeting; Meeting Abstract RECORD TYPE: Citation LANGUAGE: English

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...treatment satisfaction and effectiveness with three dosing regimens of
 intranasal fluticasone propionate (FP) in perennial nonallergic
 rhinitis (PNAR)
1997
DESCRIPTORS:
 MISCELLANEOUS TERMS: ...PERENNIAL NONALLERGIC RHINITIS ; ...
... RESPIRATORY SYSTEM DISEASE
5/3,K/29
              5:Biosis Previews(R)
DIALOG(R)File
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199799444504
0010810444
Efficacy of three different dosing regimens of fluticasone propionate (FP)
 aqueous nasal spray in the treatment of perennial nonallergic
   (PNAR)
AUTHOR: Finn A F Jr (Reprint); Howlandi W C Ii; Bronsky E A; Lumry W R;
 Pepsin P J; Rogenes P R; Westlund R; Cook C K
AUTHOR ADDRESS: Charleston, SC, USA**USA
JOURNAL: Journal of Allergy and Clinical Immunology 99 (1 PART 2): pS440
1997
    1997
CONFERENCE/MEETING: Joint Meeting of the American Academy of Allergy,
Asthma and Immunology, the American Association of Immunologists and the
Clinical Immunology Society San Francisco, California, USA February
21-26, 1997; 19970221
ISSN: 0091-6749
DOCUMENT TYPE: Meeting; Meeting Abstract
RECORD TYPE: Citation
LANGUAGE: English
...different dosing regimens of fluticasone propionate (FP) aqueous nasal
 spray in the treatment of perennial nonallergic rhinitis
1997
DESCRIPTORS:
 MISCELLANEOUS TERMS: ...PERENNIAL NONALLERGIC RHINITIS; ...
... RESPIRATORY SYSTEM DISEASE
5/3,K/30
DIALOG(R) File 5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0010810367
            BIOSIS NO.: 199799444427
Elastase in nasal lavage of cystic fibrosis and chronic fatigue syndrome
  (CFS) subjects with nonallergic rhinitis (NAR)
AUTHOR: Yuta A (Reprint); Fujita K (Reprint); Shimizu T (Reprint); Ali M;
 Clauw D; Baraniuk J N
AUTHOR ADDRESS: Mie Univ., Mie, Japan**Japan
JOURNAL: Journal of Allergy and Clinical Immunology 99 (1 PART 2): pS420
CONFERENCE/MEETING: Joint Meeting of the American Academy of Allergy,
Asthma and Immunology, the American Association of Immunologists and the
Clinical Immunology Society San Francisco, California, USA February
21-26, 1997; 19970221
ISSN: 0091-6749
DOCUMENT TYPE: Meeting; Meeting Abstract
```

RECORD TYPE: Citation LANGUAGE: English Elastase in nasal lavage of cystic fibrosis and chronic fatigue syndrome (CFS) subjects with nonallergic rhinitis (NAR) 1997 DESCRIPTORS: MISCELLANEOUS TERMS: ... NONALLERGIC RHINITIS ; RESPIRATORY SYSTEM DISEASE 5/3,K/31 DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv. 0010810366 BIOSIS NO.: 199799444426 Nonallergic rhinitis (NAR) of chronic fatigue syndrome (CFS) AUTHOR: Ali M; Gaumond E; Yuta A; Clauw D; Baraniuk J N AUTHOR ADDRESS: Georgetown Univ., Washington, DC, USA**USA JOURNAL: Journal of Allergy and Clinical Immunology 99 (1 PART 2): pS420 1997 1997 CONFERENCE/MEETING: Joint Meeting of the American Academy of Allergy, Asthma and Immunology, the American Association of Immunologists and the Clinical Immunology Society San Francisco, California, USA February 21-26, 1997; 19970221 ISSN: 0091-6749 DOCUMENT TYPE: Meeting; Meeting Abstract RECORD TYPE: Citation LANGUAGE: English Nonallergic rhinitis (NAR) of chronic fatigue syndrome (CFS) 1997 DESCRIPTORS: MISCELLANEOUS TERMS: ... NONALLERGIC RHINITIS ; RESPIRATORY SYSTEM DISEASE 5/3,K/32 DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv. BIOSIS NO.: 199799379976 Existence of c-kit receptor-positive, tryptase-negative, IgE-negative cells in human allergic nasal mucosa: A candidate for mast cell progenitor AUTHOR: Kawabori Shinichi (Reprint); Kanai Naoki; Tosho Takuro; Adachi Toshihide AUTHOR ADDRESS: Dep. Otolaryngol., Asahikawa Medical Sch., Nishikagura 4-5-3-11, Asahikawa 078, Japan**Japan JOURNAL: International Archives of Allergy and Immunology 112 (1): p36-43 1997 **1997**

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ISSN: 1018-2438

1997

... ABSTRACT: in the nasal mucosae of 11 patients with nasal allergy and of

5 patients with **nonallergic rhinitis**. From one to four of these cells in the nasal epithelium and subepithelial layer of...
DESCRIPTORS:

MISCELLANEOUS TERMS: ... RESPIRATORY SYSTEM

5/3,K/33

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0010739112 BIOSIS NO.: 199799373172

Differences in nonspecific bronchial responsiveness between patients with asthma and patients with rhinitis are not explained by type and degree of inhalant allergy

AUTHOR: Witteman Agnes M; Sjamsoedin Deman H S; Jansen Henk M; Van Der Zee Jaring S (Reprint)

AUTHOR ADDRESS: Academic Medical Cent., Univ. Amsterdam, Dep. Pulmonol., F4-239, PO Box 22700, NL-1100 DE Amsterdam, Netherlands**Netherlands
JOURNAL: International Archives of Allergy and Immunology 112 (1): p65-72
1997 1997

ISSN: 1018-2438

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

1997

... ABSTRACT: as confounding variables. In addition, a matched pair analysis was performed. Twenty-five patients with **nonallergic rhinitis** served as controls to evaluate the influence of an IgE-independent inflammatory reaction in the upper **respiratory** tract on the level of bronchial responsiveness. Furthermore, we investigated the level of nonspecific responsiveness...

...patients, the difference in level of bronchial responsiveness remained (p lt 0.001). Patients with **nonallergic rhinitis** had higher levels of nonspecific bronchial responsiveness than healthy controls and did not differ from...

DESCRIPTORS:

MISCELLANEOUS TERMS: ... RESPIRATORY SYSTEM DISEASE

5/3, K/34

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0010663876 BIOSIS NO.: 199799297936

A study of clinical and allergic aspects of rhinitis patients in Riyadh

AUTHOR: Zakzouk Siraj Mustafa; Gad-El-Rab Mohamed Osman

AUTHOR ADDRESS: Dep. ENT, King Abdulaziz Univ. Hosp., P.O Box 245, Riyadh 11411, Saudi Arabia**Saudi Arabia

JOURNAL: Annals of Saudi Medicine 16 (5): p550-553 1996 1996

ISSN: 0256-4947

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

1996

... ABSTRACT: investigated. Fifty-three (66.25%) were identified as allergic

```
and 27 (33.5%) as having nonallergic rhinitis . Medical history and
  clinical examination alone seemed to be inadequate in establishing a
  diagnosis, since...
DESCRIPTORS:
  MISCELLANEOUS TERMS: ... RESPIRATORY SYSTEM DISEASE
 5/3,K/35
DIALOG(R) File 5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0010663143 BIOSIS NO.: 199799297203
Rhinitis in Singapore
AUTHOR: Yeak S (Reprint); John A B; Chee N; Chng H H
AUTHOR ADDRESS: Dep. Otolaryngol., Tan Tock Seng Hosp., Moulmein Road,
  308433, Singapore**Singapore
JOURNAL: Allergy (Copenhagen) 51 (10): p757-758 1996 1996
ISSN: 0105-4538
DOCUMENT TYPE: Article
RECORD TYPE: Citation
LANGUAGE: English
1996
DESCRIPTORS:
 MISCELLANEOUS TERMS: ... NONALLERGIC RHINITIS ; ...
... RESPIRATORY SYSTEM...
... RESPIRATORY SYSTEM DISEASE
 5/3,K/36
DIALOG(R) File 5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199598528057
0010060224
Allergy of the upper respiratory tract
BOOK TITLE: Manual of allergy and immunology, Third edition
AUTHOR: Lierl Michelle B
BOOK AUTHOR/EDITOR: Lawlor G J (Editor); Fischer T J (Editor); Adelman D C
  (Editor)
AUTHOR ADDRESS: Univ. Cincinnati, Coll. Med., Div. Allergy/Immunol.,
 Cincinnati, OH, USA**USA
p94-111 1995
BOOK PUBLISHER: Little, Brown and Co., 34 Beacon Street, Boston,
                 Massachusetts 02108, USA
               Little, Brown and Co., London, England, UK
ISBN: 0-316-51681-3
DOCUMENT TYPE: Book; Book Chapter
RECORD TYPE: Citation
LANGUAGE: English
Allergy of the upper respiratory tract
1995
DESCRIPTORS:
  ... MAJOR CONCEPTS: Respiratory System...
... Respiration ;
 MISCELLANEOUS TERMS: ... NONALLERGIC RHINITIS WITH EOSINOPHILIA
   SYNDROME
```

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5/3,K/37
DIALOG(R) File 5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
0009545348 BIOSIS NO.: 199598013181
The usefulness of the nasal smear in understanding the relationship between
   nonallergic rhinitis with eosinophilia and recurrent purulent
  rhinitis
AUTHOR: Mansmann Paris T
AUTHOR ADDRESS: West Virginia Univ., Box 9167 Health Sci. Center,
  Morgantown, WV 26506-9167, USA**USA
JOURNAL: Pediatric Asthma Allergy and Immunology 8 (2): p117-119 1994
1994
ISSN: 0883-1874
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
The usefulness of the nasal smear in understanding the relationship between
   nonallergic rhinitis with eosinophilia and recurrent purulent
  rhinitis
 1994
ABSTRACT: Review of the literature classifies nonallergic
                                                             rhinitis with
  eosinophilia as a separate entity from purulent rhinitis, chronic
  sinusitis, and chronic hyperplastic sinusitis...
DESCRIPTORS:
  ...MAJOR CONCEPTS: Respiratory System...
... Respiration
 5/3,K/38
DIALOG(R)File
              5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
            BIOSIS NO.: 199497455830
0009434545
Treatment modalities and drugs used in vasomotor rhinitis
AUTHOR: Dainyak L B
AUTHOR ADDRESS: Sci. Cent. Audiol. Hear. Aids, Minist. Health Russ.,
  Moscow, Russia**Russia
JOURNAL: Vestnik Otorinolaringologii 0 (4): p36-41 1993 1993
ISSN: 0042-4668
DOCUMENT TYPE: Article
RECORD TYPE: Citation
LANGUAGE: Russian
1993
DESCRIPTORS:
  ...MAJOR CONCEPTS: Respiratory System...
... Respiration ;
 MISCELLANEOUS TERMS: ... NONALLERGIC
                                           RHINITIS ;
 5/3,K/39
DIALOG(R) File 5:Biosis Previews(R)
(c) 2006 BIOSIS. All rts. reserv.
```

0009383011 BIOSIS NO.: 199497404296

Synthesis of interleukin-1-alpha, interleukin-6, and interleukin-8 by cultured human nasal epithelial cells

AUTHOR: Kenney John S; Baker Coralie; Welch Mary R; Altman Leonard C (Reprint)

AUTHOR ADDRESS: Div. Allergy Infect. Dis., Dep. Med., SJ-10, Univ. Wash., Seattle, WA 98195, USA**USA

JOURNAL: Journal of Allergy and Clinical Immunology 93 (6): p1060-1067

1994 **1994** ISSN: 0091-6749

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

1994

ABSTRACT: Nasal epithelium forms the initial barrier between the environment and the **respiratory** system and may be a potential source of proinflammatory interleukins, which contribute to the pathophysiology of allergic and **nonallergic rhinitis**. To explore this possibility, epithelium and cultured human nasal epithelial cells from nasal turbinates of...

...a major source of IL-1-alpha, IL-6, and IL-8 in allergic and nonallergic rhinitis. Production of those proinflammatory cytokines by epithelial cells of the nasal and sinus mucosa may...
DESCRIPTORS:

... MAJOR CONCEPTS: Respiratory System...

... Respiration

5/3,K/40

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0009059040 BIOSIS NO.: 199497080325

Pathophysiology and pharmacotherapy of common upper respiratory diseases AUTHOR: Fireman Philip

AUTHOR ADDRESS: Children's Hospital, 3705 Fifth Ave., Pittsburgh, PA 15213, USA**USA

JOURNAL: Pharmacotherapy 13 (6 PART 2): p101S-109S 1993 1993

ISSN: 0277-0008

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

Pathophysiology and pharmacotherapy of common upper respiratory diseases 1993

ABSTRACT: The response of the upper respiratory tract to many environmental factors can be used both to analyze the body's inflammatory ...

...and its reactions often represent a major portion of the problem. The most common upper respiratory illnesses are upper respiratory infections (URIs), primarily viral, followed by secondary bacterial infections. Approximately 20% of the total population...

...is no direct allergy to hay, and there is no fever. The differential

diagnosis includes **nonallergic rhinitis** with eosinophils, which mimics the pathophysiology of allergic rhinitis but yields negative results on skin...

...adrenergic agonists are the optimum choices for congestion associated with viral URIs, and allergic or **nonallergic rhinitis**. Of major importance are accurate diagnosis, selection of appropriate therapy, and patient compliance.

DESCRIPTORS:

...MAJOR CONCEPTS: Respiratory System...

... Respiration ;

5/3,K/41

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0009003161 BIOSIS NO.: 199497024446

In vivo detection of a novel macrophage-derived protein involved in the regulation of nasal mucus-like glycoconjugate secretion

AUTHOR: Sperber Kirk (Reprint); Sylvester Clewert; Gollub Edith; Goswami Satindra; Kalb Thomas H; Druce Howard; Rutledge Joyce; Marom Zvi AUTHOR ADDRESS: Div. Clinical Immunol., Box 1089, 1 Gustave Levy Pl., New York, NY 10029, USA**USA

JOURNAL: Journal of Allergy and Clinical Immunology 92 (4): p581-588 1993

ISSN: 0091-6749

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

1993

...ABSTRACT: D-10) on frozen sections (n = 5) of nasal turbinates from patients with allergic and **nonallergic rhinitis** who were undergoing rhinoplasty and measured MMS-68 levels in nasal lavages from patients who

DESCRIPTORS:

...MAJOR CONCEPTS: Respiratory System...

... Respiration

MISCELLANEOUS TERMS: ALLERGIC VS. NONALLERGIC RHINITIS PATIENT...

5/3,K/42

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0007708908 BIOSIS NO.: 199191091799

HISTAMINE CHALLENGING TEST ON THE NASAL MUCOSA OF THE PATIENTS WITH ALLERGIC RHINITIS

AUTHOR: YANG P (Reprint); TAO Z

AUTHOR ADDRESS: DEP OTOLARYNGOL, FIRST AFFILIATED HOSP, HUNAN MED UNIV** CHINA

JOURNAL: Hunan Yike Daxue Xuebao 15 (4): p369-371 1990

ISSN: 1000-5625

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: CHINESE ABSTRACT: Ninety-six patients with perennial allergic rhinitis, 20 patients with either vasomotor or eosinophilic nonallergic rhinitis , and 33 healthy subjects were all challenged with topical application 10-3M histamine on the... **DESCRIPTORS:** ...MAJOR CONCEPTS: Respiratory System... ... Respiration ; 5/3,K/43 DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv. 0007259093 BIOSIS NO.: 199090043572 EFFICACY AND SAFETY OF INTRANASAL BUDESONIDE IN THE TREATMENT OF PERENNIAL RHINITIS IN ADULTS AND CHILDREN AUTHOR: DAY J H (Reprint); ANDERSSON C B; BRISCOE M P AUTHOR ADDRESS: DIV ALLERGY IMMUNOLOGY, KINGSTON GENERAL HOSP, KINGSTON, ONTARIO, CANADA K7L 2V7**CANADA JOURNAL: Annals of Allergy 64 (5): p445-450 1990 ISSN: 0003-4738 DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: ENGLISH 1990 ... ABSTRACT: study of 51 children (6 to 18 years) and 48 adults with perennial (allergic or nonallergic) rhinitis . The trial commenced with a 2-week baseline period without treatment for perennial rhinitis. This... DESCRIPTORS: ... MAJOR CONCEPTS: Respiratory System... ... Respiration 5/3,K/44 DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv. BIOSIS NO.: 198732090223 0005361332 SACCHARIN-BISMUTH SUBGALLATE S-BSG TEST FOR NASAL MUCOCILIARY CLEARANCE NMC AN IMPROVED METHOD AUTHOR: HUBBLE M A (Reprint); JALOWAYSKI A A; MELTZER E O; KEMP J P AUTHOR ADDRESS: SAN DIEGO, CALIF, USA**USA JOURNAL: Journal of Allergy and Clinical Immunology 79 (1): p254 1987 CONFERENCE/MEETING: FORTY-THIRD ANNUAL MEETING OF THE AMERICAN ACADEMY OF

ALLERGY AND IMMUNOLOGY, WASHINGTON, D.C., USA, FEB. 19-25, 1987. J ALLERGY

CLIN IMMUNOL. ISSN: 0091-6749

DOCUMENT TYPE: Meeting RECORD TYPE: Citation LANGUAGE: ENGLISH

DESCRIPTORS: ABSTRACT HUMAN ALLERGIC RHINITIS NONALLERGIC RHINITIS
DESCRIPTORS:

... MAJOR CONCEPTS: Respiratory System...

... Respiration

5/3,K/45

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0004800235 BIOSIS NO.: 198580109130

AN IMMUNOHISTOCHEMICAL STUDY OF PLASMA CELLS IN HUMAN ALLERGIC AND NON-ALLERGIC RHINITIS

AUTHOR: AE H J (Reprint); YOEM B W; LEE D

AUTHOR ADDRESS: DEP PATHOL, COLL MED, KOREA UNIV**SOUTH KOREA JOURNAL: Korea University Medical Journal 22 (1): p205-214 1985

ISSN: 0378-648X

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: KOREAN

1985

- ...ABSTRACT: of allergic rhinitis, was invariably noted in the clinical entity such as nasal polyposis, perennial nonallergic rhinitis and aspirin intolerance. An experimental study was performed using rabbit antihuman IgG, IgM, and IgA on the tissue sections of nasal mucosa to find differences between allergic and nonallergic rhinitis. In nonallergic rhinitis, all the cases examined showed plasma cells which react to antihuman IgG and IgA especially...
- ...the cases of allergic rhinitis. It seems to be possible to differentiate allergic rhinitis from **nonallergic rhinitis** by the scarcity of plasma cells which react antihuman IgG antibody and increase in numbers...

 DESCRIPTORS:
 - ...MAJOR CONCEPTS: Respiratory System...

... Respiration

5/3,K/46

DIALOG(R) File 5:Biosis Previews(R) (c) 2006 BIOSIS. All rts. reserv.

0002961740 BIOSIS NO.: 198069075727

ALLERGIC AND NONALLERGIC RHINITIS THEIR CHARACTERIZATION WITH ATTENTION TO THE MEANING OF NASAL EOSINOPHILIA

AUTHOR: MULLARKEY M F (Reprint); HILL J S; WEBB D R

AUTHOR ADDRESS: SECT IMMUNOL ALLERGY RHEUMATIC DIS, DEP MED, MASON CLIN,

1100 NINTH AVE, SEATTLE, WASH 98101, USA**USA

JOURNAL: Journal of Allergy and Clinical Immunology 65 (2): p122-126 1980

ISSN: 0091-6749

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: ENGLISH

ALLERGIC AND NONALLERGIC RHINITIS THEIR CHARACTERIZATION WITH ATTENTION TO THE MEANING OF NASAL EOSINOPHILIA
1980

- ABSTRACT: The differences between allergic and **nonallergic rhinitis** were examined. Patients (142) were evaluated. Forty-eight patients were diagnosed as having allergic rhinitis...
- ...no evidence for immunologic nasal disease, incriminated physical agents as precipitants, and demonstrated no associated **respiratory** pathology. These patients were classified as having vasomotor rhinitis (VMR). Twenty-one patients had symptoms...
- ...of patients with VMR, but they demonstrated nasal eosinophilia and were classified as having eosinophilic **nonallergic rhinitis** (ENR). These patients had a high prevalence of nasal polyps and were significantly more responsive...
- ...value in the evaluation of AR but provides significant information regarding therapy and prognosis in **nonallergic rhinitis**.

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\$1.06 TELNET

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